

TRANSMITTAL LETTER TO THE UNITED STATES  
DESIGNATED/ELECTED OFFICE (DO/EO/US)  
CONCERNING A FILING UNDER 35 U.S.C. 371

BIRKELUND=1

U.S. APPLICATION NO. (If known, see 37 CFR 1.5)

09/446677

INTERNATIONAL APPLICATION NO.

PCT/DK98/00266

INTERNATIONAL FILING DATE

19 June 1998

PRIORITY DATE CLAIMED

23 June 1997

TITLE OF INVENTION

SURFACE EXPOSED PROTEINS FROM CHLAMYDIA PNEUMONIAE

APPLICANT(S) FOR DO/EO/US

Svend BIRKELUND et al.

Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:

1. ☒ This is a **FIRST** submission of items concerning a filing under 35 U.S.C. 371.
2. ☐ This is a **SECOND** or **SUBSEQUENT** submission of items concerning a filing under 35 U.S.C. 371.
3. ☒ This express request to begin national examination procedures (35 U.S.C. 371(f)) at any time rather than delay examination until the expiration of the applicable time limit set in 35 U.S.C. 371(b) and PCT Articles 22 and 39(1).
4. ☒ A proper Demand for International Preliminary Examination was made by the 19th month from the earliest claimed priority date.
5. ☒ A copy of the International Application as filed (35 U.S.C. 371(c)(2))
  - a. ☐ is transmitted herewith (required only if not transmitted by the International Bureau).
  - b. ☒ has been transmitted by the International Bureau.
  - c. ☐ is not required, as the application was filed in the United States Receiving Office (RO/US).
6. ☐ A translation of the International Application into English (35 U.S.C. 371(c)(2)).
7. ☒ Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))
  - a. ☐ are transmitted herewith (required only if not transmitted by the International Bureau).
  - b. ☐ have been transmitted by the International Bureau.
  - c. ☐ have not been made; however, the time limit for making such amendments has NOT expired.
  - d. ☒ have not been made and will not be made.
8. ☐ A translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).
9. ☐ An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).
10. ☐ A translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).

Items 11. to 16. below concern document(s) or information included:

11. ☐ An Information Disclosure Statement under 37 CFR 1.97 and 1.98.
12. ☐ An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.
13. ☒ A **FIRST** preliminary amendment.  
☐ A **SECOND** or **SUBSEQUENT** preliminary amendment.
14. ☐ A substitute specification.
15. ☐ A change of power of attorney and/or address letter.
16. ☒ Other items or information:
  1. A courtesy copy of the specification as originally filed.
  2. A courtesy copy of the first page of the International Publication (WO98/58953).
  3. A courtesy copy of the International Search Report.
  4. A courtesy copy of the International Preliminary Examination Report.
  5. Formal drawings, 21 sheets, figures 1-12.

09/446677

410 REC'D PCT/PTO 23 DEC 1999

17. ☒ The following fees are submitted:**BASIC NATIONAL FEE (37 CFR 1.492 (a) (1) - (5)) :**

Neither international preliminary examination fee (37 CFR 1.482)  
nor international search fee (37 CFR 1.445(a)(2)) paid to USPTO  
and International Search Report not prepared by the EPO or JPO ..... \$970.00

International preliminary examination fee (37 CFR 1.482) not paid to  
USPTO but International Search Report prepared by the EPO or JPO ..... \$840.00

International preliminary examination fee (37 CFR 1.482) not paid to USPTO but  
international search fee (37 CFR 1.445(a)(2)) paid to USPTO ..... \$760.00

International preliminary examination fee paid to USPTO (37 CFR 1.482)  
but all claims did not satisfy provisions of PCT Article 33(1)-(4) ..... \$670.00

International preliminary examination fee paid to USPTO (37 CFR 1.482)  
and all claims satisfied provisions of PCT Article 33(1)-(4) ..... \$96.00

**ENTER APPROPRIATE BASIC FEE AMOUNT =****CALCULATIONS PTO USE ONLY**

\$ 840.00

Surcharge of \$130.00 for furnishing the oath or declaration later than ☐ 20 ☐ 30  
months from the earliest claimed priority date (37 CFR 1.492(e)).

\$

CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE
Total claims	15 - 20 =	0	X \$18.00
Independent claims	13 - 3 =	10	X \$78.00

\$ -0-

\$ 780.00

MULTIPLE DEPENDENT CLAIM(S) (if applicable)

+ \$260.00

\$

**TOTAL OF ABOVE CALCULATIONS =**

\$ 1,620.00

Reduction of 1/2 for filing by small entity, if applicable. A Small Entity Statement  
must also be filed (Note 37 CFR 1.9, 1.27, 1.28).

\$

**SUBTOTAL =**

\$ 1,620.00

Processing fee of \$130.00 for furnishing the English translation later than ☐ 20 ☐ 30  
months from the earliest claimed priority date (37 CFR 1.492(f)).

\$

**TOTAL NATIONAL FEE =**

\$ 1,620.00

Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be  
accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property +

\$

**TOTAL FEES ENCLOSED =**

\$ 1,620.00

Amount to be:  
refunded

\$

charged

\$

- a. ☒ A check in the amount of \$ 1,620.00 to cover the above fees is enclosed.
- b. ☐ Please charge my Deposit Account No. \_\_\_\_\_ in the amount of \$ \_\_\_\_\_ to cover the above fees.  
A duplicate copy of this sheet is enclosed.
- c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any  
overpayment to Deposit Account No. 02-4035. A duplicate copy of this sheet is enclosed.

**NOTE:** Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the application to pending status.

SEND ALL CORRESPONDENCE TO:

**BROWDY AND NEIMARK, P.L.L.C.**  
624 Ninth Street N.W., Suite 300  
Washington, D.C. 20001

SIGNATURE:

**Iver P. Cooper**

NAME

**28,005**

REGISTRATION NUMBER

**Date of this submission: December 23, 1999**

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:	)	Art Unit:
Svend BIRKELUND et al.	)	
IA No.: PCT/DK98/00266	)	
IA Filed: 19 June 1998	)	Washington, D.C.
U.S. App. No.:	)	
(Not Yet Assigned)	)	
National Filing Date:	)	December 23, 1999
(Not Yet Received)	)	
For: SURFACE EXPOSED PROTEINS...)	)	Docket No.: BIRKELUND=1

PRELIMINARY AMENDMENT

Honorable Commissioner of Patents and Trademarks  
Washington, D.C. 20231

Sir:

Prior to action on the merits, please amend the IPER  
claims as follows:

IN THE CLAIMS

In claim 1, replace "diagnostic test" (line 1) and  
"test" (line 2) with --method--, and delete ", such as a human,".

In claims 2 and 3, replace "Diagnostic test" with  
--Method--.

In claims 11-12, insert, at the beginning of the claim,  
--Method of claim 1, comprising--, and delete ", such as a  
human,".

Rewrite claim 13 as follows:

13 (amended). A method of immunizing a mammal against  
Chlamydia pneumoniae which comprises [Use] use of an  
immunologically effective amount of a protein with the sequence  
shown in SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:6, SEQ ID NOL:8, SEQ

ID NO:10, SEQ ID NO:12, SEQ ID NO:14, SEQ ID NO:16, SEQ ID NO:18, SEQ ID NO:20, SEQ ID NO:22, or SEQ ID NO:24, or a variant or subsequence thereof, for immunising a mammal[, such as a human,] against Chlamydia pneumoniae.

Cancel claims 14 and add claim 16:

--16. The method of claim 13 wherein the protein is in undenatured form.--

Rewrite claim 15 as follows:

15 (amended). A method of immunizing a mammal against Chlamydia pneumoniae which comprises [Use] use of an immunologically effective amount of a nucleic acid fragment with the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:13, SEQ ID NO:15, SEQ ID NO:17, SEQ ID NO:19, SEQ ID NO:21, or SEQ ID NO:23, or a variant or subsequence of said nucleotide sequence which has a sequence homology of at least 50% and wherein a subsequence of 100 nucleic acids or lower shows a homology of at least 80% with any of the mentioned nucleotide sequences encoding a protein used for effecting in vivo expression of antigens against Chlamydia pneumoniae, to immunize a mammal, by administering said nucleic acid fragment under conditions conducive to expression of said protein and subsequent immunization of said mammal by said protein [in a mammal such as a human].



REMARKS

Claims have been amended to bring them into better accord with U.S. practice.

Respectfully submitted,  
BROWDY AND NEIMARK, P.L.L.C.

Attorneys for Applicant

By:

Iver P. Cooper  
Reg. No. 28,005

624 Ninth Street, N.W.  
Washington, D.C. 20001  
Telephone: (202) 628-5197  
Facsimile: (202) 737-3528  
IPC:lms  
f:\user19\wp\n-q\birklus.pa

## NOVEL SURFACE EXPOSED PROTEINS FROM CHLAMYDIA PNEUMONIAE

The present invention relates to the identification of members of a gene family from the human respiratory pathogen *Chlamydia pneumoniae*, encoding surface exposed membrane proteins of a size of approximately 89-101 kDa and of 56-57 kDa, preferably about 89.6-100.3 kDa and about 56.1 kDa. The invention relates to the novel DNA sequences, the deduced amino acid sequences of the corresponding proteins and the use of the DNA sequences and the proteins in diagnosis of infections caused by *C. pneumoniae*, in pathology, in epidemiology, and as vaccine components.

## GENERAL BACKGROUND

*C. pneumoniae* is an obligate intracellular bacteria (Christiansen and Birkelund (1992); Grayston et al. (1986)). It has a cell wall structure as Gram negative bacteria with an outer membrane, a periplasmic space, and a cytoplasmic membrane. It is possible to purify the outer membrane from Gram negative bacteria with the detergent sarkosyl. This fraction is named the 'outer membrane complex (OMC)' (Caldwell et al. (1981)). The COMC (Chlamydia outer membrane complex) of *C. pneumoniae* contains four groups of proteins: A high molecular weight protein 98 kDa as determined by SDS-PAGE, a double band of the cysteine rich outer membrane protein 2 (Omp2) protein of 62/60 kDa, the major outer membrane protein (MOMP) of 38 kDa, and the low-molecular weight lipo-protein Omp3 of 12 kDa. The Omp2/Omp3 and MOMP proteins are present in COMC from all Chlamydia species, and these genes have been cloned from both *C. trachomatis*, *C. psittaci* and *C. pneumoniae*. However, the gene encoding 98 kDa protein from *C. pneumoniae* COMC have not been characterized or cloned.

The current state of *C. pneumoniae* serology and detection

*C. pneumoniae* is an obligate intra-cellular bacteria belonging to the genus Chlamydia which can be divided into

four species: *C. trachomatis*, *C. pneumoniae*, *C. psittaci* and *C. pecorum*. Common for the four species is their obligate intra cellular growth, and that they have a biphasic life cycle, with an extracellular infectious particle (the elementary body, EB), and an intercellular replicating form (the reticulate body, RB). In addition the Chlamydia species are characterized by a common lipopolysaccharide (LPS) epitope that is highly immunogenic in human infection. *C. trachomatis* is causing the human ocular infection (trachoma) and genital infections. *C. psittaci* is a variable group of animal pathogens where the avian strains can occasionally infect humans and give rise to a severe pneumonia (ornithosis). The first *C. pneumoniae* isolate was obtained from an eye infection, but it was classified as a non-typable Chlamydia. Under an epidemic outbreak of pneumonia in Finland it was realized that the patients had a positive reaction in the Chlamydia genus specific test, (the lygranum test), and the patients showed a titre increase to the untyped Chlamydia isolates. Similar isolates were obtained in an outbreak of upper respiratory tract infections in Seattle, and the Chlamydia isolates were classified as a new species, *Chlamydia pneumoniae* (Grayston et al. (1989)). In addition, *C. pneumoniae* is suggested to be involved in the development of atherosclerotic lesions and for initiating bronchial asthma (Kuo et al. (1995)). These two conditions are thought to be caused by either chronic infections, by a hypersensitivity reaction, or both.

#### Diagnosis of *Chlamydia pneumoniae* infections

Diagnosis of acute respiratory tract infection with *C. pneumoniae* is difficult. Cultivation of *C. pneumoniae* from patient samples is insensitive, even when proper tissue culture cells are selected for the isolation. A *C. pneumoniae* specific polymerase chain reaction (PCR) has been developed by Campbell et al. (1992).

Even though *Chlamydia pneumoniae* has in several studies been detected by this PCR it is debated whether this method is suitable for detection under all clinical situations. The reason for this is, that the cells carrying *Chlamydia pneumoniae* in acute respiratory infections have not been determined, and that a chronic carrier state is expected but it is unknown in which organs and cells they are present. Furthermore, the PCR test is difficult to perform due to the low yield of these bacteria and due to the presence of inhibitory substances in the patient samples. Therefore, it will be of great value to develop sensitive and specific sero-diagnostics for detecting both acute and chronic infections. Sero-diagnosis of *Chlamydia* infections is currently based on either genus specific tests as the Lygranum test and ELISA, measuring the antibodies to LPS, or the more species specific tests where antibodies to purified EBs are measured by microimmuno fluorescence (Micro-IF) (Wang et al. (1970)). However, the micro-IF method is read by microscopy, and in order to ensure correct readings the result must be compared to the results with *C. trachomatis* used as antigen due to the cross-reacting antibodies to the common LPS epitope. Thus, there exists in the art an urgent need for development of reliable methods for species specific diagnosis of *Chlamydia pneumoniae*, as has been expressed in Kuo et al. (1995); "...a rapid reliable laboratory test of infection for the clinical laboratory is a major need in the field". Furthermore, the possible involvement of *C. pneumoniae* in atherosclerosis and bronchial asthma clearly warrants the development of an effective vaccine.

### 30 DETAILED DISCLOSURE OF THE INVENTION

The present invention aims at providing means for efficient diagnosis of infections with *Chlamydia pneumoniae* as well as the development of effective vaccines against infection with this microorganism. The invention thus relates to species specific diagnostic tests for infection in a mammal, such as a human, with *Chlamydia pneumoniae*, said tests being based on

the detection of antibodies against surface exposed membrane proteins of a size of approximately 89-101 kDa and of 56-57 kDa, preferably of about 89.6-100.3 kDa and about 56.1 kDa (the range in size of the deduced amino acid sequences was from 100.3 to 89.6 except for Omp13 with the size of 56.1 kDa), or the detection of nucleic acid fragments encoding such proteins or variants or subsequences thereof. The invention further relates to the amino acid sequences of proteins according to the invention, to variants and subsequences thereof, and to nucleic acid fragments encoding these proteins or variants or subsequences thereof. The present invention further relates to antibodies against proteins according to the invention. The invention also relates to the use of nucleic acid fragments and proteins according to the invention in diagnosis of *Chlamydia pneumoniae* and vaccines against *Chlamydia pneumoniae*.

Prior to the disclosure of the present invention only a very limited number of genes from *C. pneumoniae* had been sequenced. These were primarily the genes encoding known *C. trachomatis* homologues: MOMP, Omp2, Omp3, Kdo-transferase, the heat shock protein genes GroEl/Es and DnaK, a ribonuclease P homologue and a gene encoding a 76 kDa protein of unknown function. The reason why so few genes have been cloned to date is the very low yield of *C. pneumoniae* which can be obtained after purification from the host cells. After such purification the DNA must be purified from the EBs, and at this step the *C. pneumoniae* DNA can easily be contaminated with host cell DNA. In addition to these inherent difficulties, it is exceedingly difficult to cultivate *C. pneumoniae* and use DNA technology to produce expression libraries with very low amounts (few  $\mu$ g) of DNA. It has been known since 1993 (Melgosa et al., 1993, that a 98 kDa protein is present in OMC from *C. pneumoniae*. Even though the protein bands of 98 kDa was mentioned to be part of the OMC of *C. pneumoniae* by Melgosa, the gene sequences and thus the deduced amino acid sequences have not been determined. Only

bands originating from *Chlamydia pneumoniae* proteins in general separated by SDS-PAGE are describe therein.

However, the gene encoding this protein has not been determined before the present invention. Only a very weak or no reaction with patient sera can be observed to the 98 kDa protein (Campbell et al. 1990) and prior to the work of the present inventors it has not been recognized that the 89-101 kDa proteins are surface exposed or that they in fact is immunogenic (see below). In this report it is described that a number of human serum samples reacts with a *C. pneumoniae* protein that in SDS-PAGE migrate as 98 kDa. The protein was not further characterized and it is therefore not in conflict with the present application.

Campbell et al. (1990) described that sera from four patients from which *Chlamydia pneumoniae* was isolated reacted with bands of 98 kDa in immunoblotting using whole-cell lysates. They also showed that no proteins with similar molecular weights were recognised by serum samples in either *Chlamydia trachomatis* or *Chlamydia psittaci* and they therefore suggest that the protein present in the 98 kDa band could be used as a potential diagnostic tool for the recognition of *Chlamydia pneumoniae* infection. The protein content within the 98 kDa region was not further characterised and its localisation within the *Chlamydia* was not shown.

Halme et al. (1997) described the presence of human T-cell epitopes in *C. pneumoniae* proteins of 92-98 kDa. The proteins were eluted from SDS-PAGE of total chlamydia proteins but the identity of the proteins were not determined.

Use of antibodies to screen expression libraries is a well known method to clone fragments of genes encoding antigenic parts of proteins. However, since patient sera do not show a significant reaction with the 98 kDa protein it has not been possible to use patient serum to clone the proteins.

It was known that monoclonal antibodies generated by the inventors reacted with conformational epitopes on the surface of *C. pneumoniae* and that they also reacted with *C. pneumoniae* OMC by immuno-electron microscopy (Christiansen et al. 1994). Furthermore, the 98 kDa protein is the only unknown protein from the *C. pneumoniae* OMC (Meigosa et al. 1993). The present inventors chose to take an unconventional step in order

AMENDED SHEET

5a

to clone the gene encoding the hitherto unknown 98 kDa protein: *C. pneumoniae* OMC was purified and the highly immunogenic conformational epitopes were destroyed by SDS-treatment of the antigen before immunization. Thereby an antibody (PAB 150) to less immunogenic linear epitopes was obtained. This provided the possibility to obtain an

5

AMENDED SHEET

antiserum which could detect the protein, and it was shown that a gene family encoding the 89-101 kDa and 56 proteins according to the invention could be detected in colony blotting of recombinant *E. coli*.

5 Mice infected with *C. pneumoniae* generate antibodies to the proteins identified by the inventors and named Omp4-15, but do not recognize the SDS treated heat denatured antigens normally used for SDS-PAGE and immunoblotting. However, a strong reaction was seen if the antigen was not heat  
10 denatured. It is therefore highly likely that if a similar reaction is seen in connection with human infections the antigens of the present invention will be of invaluable use in sero-diagnostic tests and may very likely be used as a vaccine for the prevention of infections.

15 By generating antibodies against COMC from *C. pneumoniae* a polyclonal antibody (PAB 150) was obtained which reacted with all the proteins. This antibody was used to identify the genes encoding the 89.6-101.3 kDa and 56.1 kDa proteins in an  
20 expression library of *C. pneumoniae* DNA. A problem in connection with the present invention was that a family comprising a number of similar genes were found in *C. pneumoniae*. Therefore, a large number of different clones were required to identify clusters of fragments. Only because  
25 the rabbit antibody generated by the use of SDS-denatured antigens contained antibodies to a high number of different epitopes positioned on different members of the protein family did the inventors succeed in cloning and sequencing four of the genes. One gene was fully sequenced, a second was  
30 sequenced except for the distal part and shorter fragments of two additional genes were obtained by this procedure. To obtain the DNA sequence of the additional genes and to search for more members of the gene family long range PCR with primers derived from the sequenced genes, and primers from  
35 the genes already published in the database were used. This approach gave rise to the detection of additional eight genes belonging to this family. The genes were situated in two gene



clusters: Omp12,11,10,5,4,13 and 14 in one cluster and Omp6,7,8,9 and 15 in the second. Full sequence was obtained from Omp4,5,6,7,8,9,10,11 and 13, and partial sequence of Omp12,14. Omp13 was a truncated gene of 1545 nucleotides. The rest of the full length genes were from 2526 (Omp7) to 2838 (Omp15) nucleotides. The deduced amino acid sequences revealed putative polypeptides of 89.6 to 100.3 kDa, except for Omp13 of 56.1 kDa. Alignment of the deduced amino acid sequences showed a maximum identity of 49% (Omp5/Omp9) when all the sequences were compared. Except for Omp13, the lowest homology was to Omp7 with no more than 34% identity to any of the other amino acid sequences. The scores for Omp13 was from 29-32% to all the other sequences.

In the present context SEQ ID Nos. 1 and 2 correspond to Omp4, SEQ ID Nos 3 and 4 correspond to Omp5, SEQ ID Nos 5 and 6 correspond to Omp6, SEQ ID Nos 7 and 8 correspond to Omp7, SEQ ID Nos 9 and 10 correspond to Omp8, SEQ ID Nos 11 and 12 correspond to Omp9, SEQ ID Nos 13 and 14 corresponds to Omp10, SEQ ID Nos 15 and 16 corresponds to Omp11, SEQ ID Nos 17 and 18 corresponds to Omp12, SEQ ID Nos 19 and 20 corresponds to Omp13; SEQ ID Nos 21 and 22 corresponds to Omp14, and SEQ ID Nos 23 and 24 corresponds to Omp15.

The estimated size of the Omp proteins of the of the present invention are listed in the following. Omp 4 has a size of 98.9 kDa, Omp5 has an estimated size of 97.2 kDa, Omp6 has an estimated size of 100.3 kDa, Omp7 has an estimated size of 89.7 kDa, Omp8 has an estimated size of 90.0 kDa, Omp9 has an estimated size of 96.7 kDa, Omp10 has an estimated size of 98.4 kDa, Omp11 has an estimated size of 97.6 kDa, Omp13 has an estimated size of 56.1 kDa, Omp 12 and 14 being partial.

Furthermore, SEQ ID No 25 is a subsequence of SEQ ID No 3, SEQ ID No 26 is a subsequence of SEQ ID No 4, SEQ ID No 27 is a subsequence of SEQ ID No 5, SEQ ID No 28 is a subsequence of SEQ ID No 6, SEQ ID No 29 is a subsequence of SEQ ID No 7, and SEQ ID No 30 is a subsequence of SEQ ID No 8.

Part of the omp proteins were expressed as fusion proteins, and mice polyclonal monospecific antibodies against the proteins were produced. The antibodies reacted with the surface of *C. pneumoniae* in both immunofluorescence and immunoelectron microscopy. This shows for the first time that the 89-101 kDa and 56-57 kDa protein family in *C. pneumoniae* comprises surface exposed outer membrane proteins. This important finding leads to the realization that members of the 89-101 kDa and 56-57 kDa *C. pneumoniae* protein family are good candidates for the development of a sero diagnostic test for *C. pneumoniae*, as well as the development of a vaccine against infections with *C. pneumoniae* based on using these proteins. Furthermore, the proteins may be used as epidemiological markers, and polyclonal monospecific sera against the proteins can be used to detect *C. pneumoniae* in human tissue or detect *C. pneumoniae* isolates in tissue culture. Also, the genes encoding the 89-101 kDa and 56-57 kDa such as the 89.6-100.3 kDa and 56.1 protein family may be used for the development of a species specific diagnostic test based on nucleic acid detection/amplification.

The full length Omp4 was cloned into an expression vector system that allowed expression of the Omp4 polypeptide. This polypeptide was used as antigen for immunization of a rabbit. Since the protein was purified under denaturing condition the antibody did not react with the native surface of *C. pneumoniae*, but it reacted with a 98 kDa protein in immunoblotting where purified *C. pneumoniae* EB was used as antigen. Furthermore, the antibody reacted in paraffin embedded sections of lung tissue from experimentally infected mice.

A broad aspect of the present invention relates to a species specific diagnostic test for infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said test comprising detecting in a patient or preferable in a patient sample the presence of antibodies against proteins from the outer membrane of *Chlamydia pneumoniae*, said proteins being of a

molecular weight of 89-101 kDa or 56-57 kDa, or detecting the presence of nucleic acid fragments encoding said outer membrane proteins or fragments thereof.

5 In the context of the present application, the term "patient sample" should be taken to mean an amount of serum from a patient, such as a human patient, or an amount of plasma from said patient, or an amount of mucosa from said patient, or an amount of tissue from said patient, or an amount of  
10 expectorate, forced sputum or a bronchial aspirate, an amount of urine from said patient, or an amount of cerebrospinal fluid from said patient, or an amount of atherosclerotic lesion from said patient, or an amount of mucosal swaps from said patient, or an amount of cells from a tissue culture  
15 originating from said patient, or an amount of material which in any way originates from said patient. The in vivo test in a human according to the present invention includes a skin test known in the art such as an intradermal test, e.g. similar to a Mantoux test. In certain patients being very  
20 sensitive to the test, such as is often the case with children, the test could be non-invasive, such as a superficial test on the skin, e.g. by use of a plaster

In the present context, the term 89-101 kDa protein means proteins normally present in the outer membrane of *Chlamydia pneumoniae*, which in SDS-PAGE can be observed as one or more  
25 bands with an apparent molecular weight substantially in the range of 89-101 kDa. From the deduced amino acid sequences the molecular size varies from 89.6 to 100.3 kDa.

Within the scope of the present invention are species  
30 specific sero-diagnostic tests based on the usage of the genes belonging to the gene family disclosed in the present application.

Preferred embodiments of the present invention relate to species specific diagnostic tests according to the invention,  
35 wherein the outer membrane proteins have sequences selected

from the group consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24.

- 5 When used in connection with proteins according to the present invention the term "variant" should be understood as a sequence of amino acids which shows a sequence similarity of less than 100% to one of the proteins of the invention. A variant sequence can be of the same size or it can be of a different size as the sequence it is compared to. A variant will typically show a sequence similarity of preferably at least 50%, preferably at least 60%, more preferably at least 70%, such as at least 80%, e.g. at least 90%, 95% or 98%.

- 15 The term "sequence similarity" in connection with sequences of proteins of the invention means the percentage of identical and conservatively changed amino acid residues (with respect to both position and type) in the proteins of the invention and an aligned protein of equal or different length. The term "sequence identity" in connection with sequences of proteins of the invention means the percentage of identical amino acid with respect to both position and type in the proteins of the invention and an aligned protein of equal or different length.

- 25 Within the scope of the present invention are subsequences of one of the proteins of the invention, meaning a consecutive stretch of amino acid residues taken from SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24. A subsequence will typically comprise at least 100 amino acids, preferably at least 80 amino acids, more preferably at least 70 amino acids, such as 50 amino acids. It might even be as small as 10-50 amino acids, such as 20-40 amino acids, e.g. about 30 amino acids. A subsequence will typically show a sequence  
35 homology of at least 50%, preferably at least 60%, more

preferably at least 70%, such as at least 80%, e.g. at least 90%, 95% or 98%.

Diagnostic tests according to the invention include immunoassays selected from the group consisting of a direct or indirect EIA such as an ELISA, an immunoblot technique such as a Western blot, a radio immuno assay, and any other non-enzyme linked antibody binding assay or procedure such as a fluorescence, agglutination or precipitation reaction, and nephelometry.

10 A preferred embodiment of the present invention relates to species specific diagnostic tests according to the invention, said test comprising an ELISA, wherein antibodies against the proteins of the invention or fragments thereof are detected in samples.

15 A preferred embodiment of the invention, is an ELISA based on detection in samples of antibodies against proteins of the invention. The ELISA may use proteins of the invention, or variants thereof, i.e. the antigen, as coating agent. An ELISA will typically be developed according to standard  
20 methods well known in the art, such as methods described in "Antibodies; a laboratory manual", Ed. David Lane Harlow, Cold Spring Harbor laboratories (1988), which is hereby incorporated by reference.

Recombinant proteins will be produced using DNA sequences  
25 obtained essentially using methods described in the examples below. Such DNA sequences, comprising the entire coding region of each gene in the gene family of the invention, will be cloned into an expression vector from which the deduced protein sequence can be purified. The purified proteins will  
30 be analyzed for reactivity in ELISA using both monoclonal and polyclonal antibodies as well as sera from experimentally infected mice and human patient sera.

From the experimentally infected mice sera it is known that non-linear epitopes are recognized predominantly. Thus, it is contemplated that different forms of purification schemes known in the art will be used to analyze for the presence of discontinuous epitopes, and to analyze whether the human immune response is also directed against such epitopes.

Preferred embodiments of the present invention relate to species specific diagnostic tests according to the invention, wherein the nucleic acid fragments have sequences selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, and SEQ ID NO: 23.

In connection with nucleic acid fragments according to the present invention the term "variant" should be understood as a sequence of nucleic acids which shows a sequence homology of less than 100%. A variant sequence can be of the same size or it can be of a different size as the sequence it is compared to. A variant will typically show a sequence homology of at least 50%, preferably at least 60%, more preferably at least 70%, such as at least 80%, e.g. at least 90%, 95% or 98%.

The term "sequence homology" in connection with nucleic acid fragments of the invention means the percentage of matching nucleic acids (with respect to both position and type) in the nucleic acid fragments of the invention and an aligned nucleic acid fragment of equal or different length.

In order to obtain information concerning the general distribution of each of the genes according to the present invention, PCR will be performed for each gene on all available *C. pneumoniae* isolates. This will provide information on the general variability of the genes or nucleic acid fragments of the invention. Variable regions will be sequenced. From patient samples PCR will be used to

amplify variable parts of the genes for epidemiology. Non-variable parts will be used for amplification by PCR and analyzed for possible use as a diagnostic test. It is contemplated that if variability is discovered, PCR of  
5 variable regions can be used for epidemiology. PCR of non-variable regions can be used as a species specific diagnostic test. Using genes encoding proteins known to be invariable in all known isolates prepared as targets for PCR to genes encoding proteins with unknown function.

10 Particularly preferred embodiments of the present invention, relate to diagnostic tests according to the invention, wherein detection of nucleic acid fragments is obtained by using nucleic acid amplification, preferably polymerase chain reaction (PCR).

15 Within the scope of the present invention is a PCR based test directed at detecting nucleic acid fragments of the invention or variants thereof. A PCR test will typically be developed according to methods well known in the art and will typically comprise a PCR test capable of detecting and differentiating  
20 between nucleic acid fragments of the invention. Preferred are quantitative competitive PCR tests or nested PCR tests. The PCR test according to the invention will typically be developed according to methods described in detail in EP B 540 588, EP A 586 112, EP A 643 140 OR EP A 669 401, which  
25 are hereby incorporated by reference.

Within the scope of the present invention are variants and subsequences of one of the nucleic acid fragments of the invention, meaning a consecutive stretch of nucleic acids taken from SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID  
30 NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 19, SEQ ID NO: 21, or SEQ ID NO: 23. A variant or subsequence will preferably comprise at least 100 nucleic acids, preferably at least 80 nucleic acids, more preferably at least 70 nucleic acids, such as at least 50 nucleic acids.  
35 It might even be as small as 10-50 nucleic acids, such as

20-40 nucleic acids, e.g. about 30 nucleic acids. A  
subsequence will typically show a sequence homology of at  
least 30%, preferably at least 60%, more preferably at least  
70%, such as at least 80%, e.g. at least 90%, 95% or 98%. The  
5 shorter the subsequence, the higher the required homology.  
Accordingly, a subsequence of 100 nucleic acids or lower must  
show a homology of at least 80%.

A very important aspect of the present invention relates to  
proteins of the invention derived from *Chlamydia pneumoniae*  
10 having amino acid sequences selected from the group  
consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ  
ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID  
NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ  
ID NO: 24 having a sequence similarity of at least 50%,  
15 preferably at least 60%, more preferably at least 70%, such  
as at least 80%, e.g. at least 90%, 95% or 98% and a similar  
biological function.

By the term "similar biological function" is meant that the  
protein shows characteristics similar with the proteins  
20 derivable from the membrane proteins of *Chlamydia pneumoniae*.  
Such proteins comprise repeated motifs of GGAI (at least 2,  
preferable at least 3 repeats) and/or conserved positions of  
tryptophan, (w).

Comparison of the DNA sequences from genes encoding Omp4-15  
25 shows that the overall similarity between the individual  
genes ranges between 43-55%. Comparison of the amino acid  
sequences of Omp4-15 shows 34-49% identity and 53-64%  
similarity. The homology is generally scattered along the  
entire length of the deduced amino acids. However, as seen  
30 from figure 8 A - J there are some regions in which the  
homology is more pronounced. This is seen in the repeated  
sequence where the sequence GGAI is repeated 4-7 times in the  
genes. It is interesting that the DNA homology is not  
conserved for the sequences encoding the four amino acids  
35 GGAI. This may indicate a functional role of this part of the



protein and indicates that the repeated structure did not occur by a duplication of the gene. In addition to the four amino acid repeats GGAI a region from amino acid 400 to 490 has a higher degree of homology than the rest of the protein, with the conserved sequence FYDPI occurring in all sequences. As further indication of similarity in function the amino acid tryptophan (W) is perfectly conserved at 4-6 localizations in the C-terminal part of the protein.

Since none of the genes and deduced amino acid sequences of the invention are identical the following is within the scope of the present invention; production of monospecific antibodies, the use of said antibodies for characterizing which *C. pneumoniae* proteins are expressed, the use of said antibodies for characterizing at which time during developmental life cycle said *C. pneumoniae* proteins are expressed, and the use of said antibodies for characterizing the precise cellular localization of said *C. pneumoniae* proteins. Also within the scope of the present invention is the use of monospecific antibodies against proteins of the invention for determining which part of said proteins is surface exposed and how proteins in the *C. pneumoniae* COMC interact with each other.

Preferred embodiments of the present invention relate to polypeptides which comprise subsequences of the proteins of the invention, said subsequences comprising the sequence GGAI. Further preferred embodiments of the present invention relate to polypeptides which comprise subsequences of the proteins of the invention, said subsequences comprising the sequence FSGE.

Polypeptides according to the invention will typically be of a length of at least 6 amino acids, preferably at least 15 amino acids, preferably at least 20 amino acids, preferably at least 25 amino acids, preferably at least 30 amino acids, preferably at least 35 amino acids, preferably at least 40 amino acids, preferably at least 45 amino acids, preferably

at least 50 amino acids, preferably at least 55 amino acids, preferably at least 100 amino acids.

A very important aspect of the present invention relates to nucleic acid fragments of the invention derived from

5 *Chlamydia pneumoniae*, variants and subsequences thereof.

Another important aspect of the present invention relates to antibodies against the proteins according to the invention, such antibodies including polyclonal monospecific antibodies and monoclonal antibodies against proteins with sequences

10 selected from the group consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24.

A very important aspect of the present invention relates to  
15 diagnostic kits for the diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said kits comprising one or more proteins with amino acid sequences selected from the group consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24.

Another very important aspect of the present invention relates to diagnostic kits for the diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said  
25 kits comprising antibodies against a protein with an amino acid sequence selected from the group consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24.  
30 Antibodies included in a diagnostic kit according to the invention can be polyclonal or monoclonal or a mixture hereof.

Still another very important aspect of the present invention relates to diagnostic kits for the diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said kits comprising one or more nucleic acid fragments with

5 sequences selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, and SEQ ID NO: 23.

10 An aspect of the present invention relates to a composition for immunizing a mammal, such as a human, against *Chlamydia pneumoniae*, said composition comprising one or more proteins with amino acid sequences selected from the group consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16,  
15 SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24.

An important role for the proteins of the invention in prevention of infection of a mammal, such as a human, with *C. pneumoniae* is expected. Thus proteins of the invention,  
20 including variants and subsequences will be produced, typically by using recombinant techniques, and will then be used as an antigen in immunization of mammals, such as rabbits. Subsequently, the hyper immune sera obtained by the immunization will be analyzed for protection against *C.*  
25 *pneumoniae* infection using a tissue culture assay. In addition it is contemplated that monoclonal antibodies will be produced, typically using standard hybridoma techniques, and analyzed for protection against infection with *C. pneumoniae*.

30 It is envisioned that particularly interesting and immunogenic epitopes will be found in connection with the proteins of the invention, which will comprise subsequences of said proteins. It is preferred to use polypeptides comprising such subsequences of the proteins of the invention

in immunizing a mammal, such as a human, against *Chlamydia pneumoniae*.

An important aspect of the present invention relates to the use of proteins with sequences selected from the group

- 5 consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24 in diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*.

- 10 A preferred embodiment of the present invention relates to the use of proteins according to the invention in an undenatured form, in diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*.

- A very important aspect of the present invention relates to
- 15 the use of proteins with sequences selected from the group consisting of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, and SEQ ID NO: 24, for immunizing a mammal, such as a human, against
- 20 *Chlamydia pneumoniae*.

A preferred embodiment of the present invention relates to the use of proteins according to the invention in an undenatured form, for immunizing a mammal, such as a human, against *Chlamydia pneumoniae*.

- 25 A very important aspect of the present invention relates to the use of nucleic acid fragments with nucleotide sequences selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, and SEQ ID NO: 23 for immunizing a mammal,
- 30 such as a human, against *Chlamydia pneumoniae*.

It is envisioned that one type of vaccine against *C. pneumoniae* will be developed by using gene-gun vaccination of mice. Typically, different genetic constructs containing nucleic acid fragments, combinations of nucleic acid

5 fragments according to the invention will be used in the gene-gun approach. The mice will then subsequently be analyzed for production of both humoral and cellular immune response and for protection against infection with *C. pneumoniae* after challenge herewith.

10 In line with this, the invention also relates to the uses of the proteins of the invention as a pharmaceutical (a vaccine) as well as to the uses thereof for the preparation of a vaccine against infections with *Chlamydia pneumoniae*.

Preparation of vaccines which contain protein sequences as  
15 active ingredients is generally well understood in the art, as exemplified by U.S. Patents 4,608,251; 4,601,903; 4,599,231; 4,599,230; 4,596,792; and 4,578,770, all incorporated herein by reference. Typically, such vaccines are prepared as injectables either as liquid solutions or suspen-  
20 sions; solid forms suitable for solution in, or suspension in, liquid prior to injection may also be prepared. The preparation may also be emulsified. The active immunogenic ingredient is often mixed with excipients which are pharmaceutically acceptable and compatible with the active ingredi-  
25 ent. Suitable excipients are, for example, water, saline, dextrose, glycerol, ethanol, or the like, and combinations thereof. In addition, if desired, the vaccine may contain minor amounts of auxiliary substances such as wetting or emulsifying agents, pH buffering agents, or adjuvants which  
30 enhance the effectiveness of the vaccines.

The vaccines are conventionally administered parenterally, by injection, for example, either subcutaneously or intramuscularly. Additional formulations which are suitable for other modes of administration include suppositories and, in some  
35 cases, oral formulations. These compositions take the form of

solutions, suspensions, tablets, pills, capsules, sustained release formulations or powders and contain 10-95% of active ingredient, preferably 25-70%, and optionally a suitable carrier.

- 5 The protein sequences may be formulated into the vaccine as neutral or salt forms known in the art. The vaccines are administered in a manner compatible with the dosage formulation, and in such amount as will be therapeutically effective and immunogenic. The quantity to be administered  
10 depends on the subject to be treated. Suitable dosage ranges are of the order of several hundred micrograms active ingredient per vaccination with a preferred range from about 0.1  $\mu\text{g}$  to 1000  $\mu\text{g}$ . The immune response may be enhanced if the vaccine further comprises an adjuvant substance as known in  
15 the art. Other possibilities involve the use of immunomodulating substances such as lymphokines (e.g. IFN- $\gamma$ , IL-2 and IL-12) or synthetic IFN- $\gamma$  inducers such as poly I:C in combination with the above-mentioned adjuvants.

- It is also possible to produce a living vaccine by introducing, into a non-pathogenic microorganism, at least one  
20 nucleic acid fragment encoding a protein fragment or protein of the invention, and effecting expression of the protein fragment or the protein on the surface of the microorganism. (e.g. in the form of a fusion protein including a membrane  
25 anchoring part or in the form of a slightly modified protein or protein fragment carrying a lipidation signal which allows anchoring in the membrane). The skilled person will know how to adapt relevant expression systems for this purpose.

- Another part of the invention is based on the fact that  
30 recent research have revealed that a DNA fragment cloned in a vector, which is non-replicative in eukaryotic cells may be introduced into an animal (including a human being) by e.g. intramuscular injection or percutaneous administration (the so-called "gene gun" approach). The DNA is taken up by e.g.  
35 muscle cells and the gene of interest is expressed by a

promoter which is functioning in eukaryotes, e.g. a viral promoter, and the gene product thereafter stimulates the immune system. These newly discovered methods are reviewed in Ulmer et al., 1993, which hereby is included by reference.

- 5 Thus, a nucleic acid fragment encoding a protein or protein of the invention may be used for effecting *in vivo* expression of antigens, i.e. the nucleic acid fragments may be used in so-called DNA vaccines. Hence, the invention also relates to a vaccine comprising a nucleic acid fragment encoding a  
10 protein fragment or a protein of the invention, the vaccine effecting *in vivo* expression of antigen by an mammal, such as a human, to whom the vaccine has been administered, the amount of expressed antigen being effective to confer substantially increased resistance to infections with  
15 *Chlamydia pneumoniae* in an mammal, such as a human.

- The efficacy of such a "DNA vaccine" can possibly be enhanced by administering the gene encoding the expression product together with a DNA fragment encoding a protein which has the capability of modulating an immune response. For instance, a  
20 gene encoding lymphokine precursors or lymphokines (e.g. IFN- $\gamma$ , IL-2, or IL-12) could be administered together with the gene encoding the immunogenic protein fragment or protein, either by administering two separate DNA fragments or by administering both DNA fragments included in the same vector.  
25 It is also a possibility to administer DNA fragments comprising a multitude of nucleotide sequences which each encode relevant epitopes of the protein fragments and proteins disclosed herein so as to effect a continuous sensitization of the immune system with a broad spectrum of these epitopes.  
30 The following experimental non-limiting examples are intended to illustrate certain features and embodiments of the invention.

## LEGENDS TO FIGURES

Figure 1. The figure shows electron microscopy of negative stained purified *C. pneumoniae* EB (A) and purified OMC (B).

Figure 2. The figure shows silver stained 15% SDS-PAGE of purified EB and OMC. Lane 1, purified *C. pneumoniae* EB; lane 2, *C. pneumoniae* OMC; lane 3, purified *C. trachomatis* EB; and lane 4 *C. trachomatis* OMC.

Figure 3. The figure shows immunoblotting of *C. pneumoniae* EB separated by 10% SDS-PAGE, transferred to nitrocellulose and reacted with rabbit anti *C. pneumoniae* OMC.

Figure 4. The figure shows coomassie blue stained 7.5% SDS-PAGE of recombinant pEX that were detected by the rabbit anti *C. pneumoniae* serum. Arrow indicated the localization of the 117 kDa b-galactosidase protein.

Figure 5. The figure shows immunoblotting of recombinant pEX clones detected by colony blotting separated by 7.5% SDS-PAGE and transferred to nitrocellulose and reacted with rabbit anti *C. pneumoniae* OMC. Lane 1, seablue molecular weight standard. Lane 2-6 pEX clones cultivated at 42°C to induce the production of the b-galactosidase fusion proteins.

Figure 6. The figure shows sequence strategy for Omp4 and Omp5. Arrows indicates primers used for sequencing.

Figure 7. *C. pneumoniae* omp genes. The genes are arranged in two clusters. In cluster 1 Omp12, 11, 10, 5, 4, 13, and 14 are found. In cluster 2 are found Omp6, 7, 8, 9, and 15.

Figure 8 A - J. The figure shows alignment of *C. pneumoniae* Omp4-15, using the program pileup in the GCG package.

Figure 9. The figure shows immunofluorescence of *C. pneumoniae* infected HeLa, 72 hrs. after infection, reacted



with mouse monospecific anti-serum against pEX3-36 fusion protein. pEX3-36 is a part of the Omp5 gene.

5 Figure 10. The figure shows immunoblotting of *C. pneumoniae* EB, lane 1-3 heated to 100°C in SDS-sample buffer, lane 4-6 unheated. Lane 1 reacted with rabbit anti *C. pneumoniae* OMC; lane 2 and 4 pre-serum; lane 3 and 5 polyclonal rabbit anti pEX1-1 fusion protein; lane 6 MAb 26.1.

10 Figure 11. The figure shows immunoblotting of *C. pneumoniae* EB, lane 1-4 heated to 100°C in SDS-sample buffer, lane 5-6 unheated. Reacted with serum from C57-black mice 14 days after infection with  $10^7$  CFU of *C. pneumoniae*. Lane 1 and 5 mouse 1; lane 2 and 6 mouse 2; lane 3 and 5 mouse 3; and lane 4 and 8 mouse 4.

15 Figure 12. The figure shows immunohistochemistry analysis of mouse lung tissue with *C. pneumoniae* inclusions present both in the bronchial epithelium and in the lung parenchyma (arrows).

## EXAMPLE 1

Cloning of the genes encoding the 98/95 kDa *C. pneumoniae* COMC proteinsPurification of *C. pneumoniae* EBs and COMC

- 5 *C. pneumoniae* was cultivated in HeLa cells. Cultivation was done according to the specifications of Miyashita and Matsumoto (1992), with the modification that centrifugation of supernatant and of the later precipitate and turbid bottom layer was carried out at 100,000 X g. The microorganism
- 10 attached to the HeLa cells by 30 minutes of centrifugation at 1000 x g, after which the cells were incubated in RPMI 1640 medium (Gibco BRL, Germany cat No. 51800-27), containing 5% foetal calf serum (FCS, Gibco BRL, Germany Cat No. 10106.169) gentamicin for two hours at 37°C in 5% CO<sub>2</sub> atmosphere. The
- 15 medium was changed to medium that in addition contained 1 mg per ml of cycloheximide. After 48 hours of incubation a coverslip was removed from the cultures and the inclusion was tested with an antibody specific for *C. pneumoniae* (MAb 26.1) (Christiansen et al. 1994) and a monoclonal antibody specific
- 20 for the species *C. trachomatis* (MAb 32.3, Loke diagnostics, Århus Denmark) to ensure that no contamination with *C. trachomatis* had occurred. The HeLa cells were tested by Hoechst stain for Mycoplasma contamination as well as by culture in BEa and BEg medium (Freund et al., 1979). Also the
- 25 *C. pneumoniae* stocks were also tested for Mycoplasma contamination by cultivation in BEa and BEg medium. No contamination with *C. trachomatis*, Mycoplasmas or bacteria were detected in cultures or cells. 72 hours post-infection the monolayer was washed in PBS, the cells were loosened in
- 30 PBS with a rubber policeman, and the Chlamydia were liberated from the host cell by sonication. The *C. pneumoniae* EBs and RBs were purified on discontinuous density gradients (Miyashita et al. (1992)). The purity of the Chlamydia EBs were verified by negative staining and electronmicroscopy
- 35 (Figure 1), only particles of a size of 0.3 to 0.5 µm were

detected in agreement with the structure of *C. pneumonia* EBs. The purified Chlamydia EBs were subjected to sarkosyl extraction as described by Caldwell et al (1981) with the modification that a brief sonication was used to suspend the COMC. The purified COMC was tested by electronmicroscopy and negative staining (Figure 1), where a folded outer membrane complex was seen.

#### SDS-PAGE analysis of purified EBs and COMC

The proteins from purified EBs and *C. pneumoniae* OMC were separated on 15% SDS-polyacrylamide gel, and the gel was silver stained (Figure 2), in lane 1 it is seen that the purified EBs contain major proteins of 100/95 kDa and a protein of 38 kDa, in the purified COMC (lane 2) these two protein groups are also dominant. In addition, proteins with a molecular weight of 62/60 kDa, 55 kDa, and 12 kDa have been enriched in the COMC preparation. When the purified *C. pneumoniae* EBs are compared to purified *C. trachomatis* EB (lane 3) it is seen that predominant protein in the *C. trachomatis* EB is the major outer membrane protein (MOMP), and it is also the dominant band in the COMC preparation of *C. trachomatis* (lane 4), and Omp2 of 60/62 kDa as well as Omp3 at 12 kDa are seen in the preparation. However, no major bands with a size of 100/95 kDa are detected as in the *C. pneumoniae* COMC preparation.

#### 25 Production of rabbit polyclonal antibodies against *C. pneumoniae* COMC

To ensure production of rabbit antibodies that would recognize all the *C. pneumoniae* proteins in immuno-blotting and colony-blotting 10 µg of COMC antigen was dissolved in 20 µl of SDS sample buffer and thereafter divided into 5 vials. The dissolved antigen was further diluted in one ml of PBS and one ml of Freund incomplete adjuvant (Difco laboratories, USA cat. No. 0639-60-6) and injected into the quadriceps muscle of a New Zealand white rabbit. The rabbit was given

three times intramuscular injections at an interval of one week, and after further three weeks the dissolved COMC protein, diluted in one ml PBS was injected intravenously, and the procedure was repeated two weeks later. Eleven weeks  
5 after the beginning of the immunization, the serum was obtained from the rabbit. Purified *C. pneumoniae* EBs were separated by SDS-PAGE, and the proteins were electrotransferred to nitrocellulose membrane. The membrane was blocked and immunostained with the polyclonal COMC  
10 antibody (Figure 3). The serum recognized proteins with a size of 100/95, 60 and 38 kDa in the EB preparation. This is in agreement with the sizes of the outer membrane proteins.

#### Cloning of the COMC proteins

Due to the cultivation of *C. pneumoniae* in HeLa cells,  
15 contaminating host cell DNA could be present in the EB preparations. Therefore, the purified EB preparations were treated with DNase to remove contaminating DNA. The *C. pneumoniae* DNA was then purified by CsCl gradient centrifugation. The *C. pneumoniae* DNA was partially digested  
20 with Sau3A and the fractions containing DNA fragments with a size of approx. 0.5 to 4.0 kb were cloned into the expression vector system pEX (Boehringer, Germany cat. No. 1034 766, 1034 774, 1034 782). The pEX vector system has a  
25  $\beta$ -galactosidase gene with multiple cloning sites in the 3' end of the  $\beta$ -galactosidase gene. Expression of the gene is regulated by the PR promoter, so the protein expression can be induced by elevating the temperature from 32 to 42°C. The colonies of recombinant bacteria were transferred to nitrocellulose membranes, and the temperature was increased  
30 to 42°C for two hours. The bacteria were lysed by placing the nitrocellulose membranes on filters soaked in 5% SDS. The colonies expressing outer membrane proteins were detected with the polyclonal antibody raised against *C. pneumoniae* COMC. The positive clones were cultivated in suspension and  
35 induced at 42°C for two hours. The protein profile of the clones were analysed by SDS-PAGE, and increases in the size

of the induced b-galactosidase were observed (Figure 4). In addition, the proteins were electrotransferred to nitrocellulose membranes, and the reaction with the polyclonal serum against COMC was confirmed (Figure 5).

## 5 Sequencing of positive COMC clones

To characterize the pEX clones, the inserted *C. pneumoniae* DNA was sequenced. The resulting DNA sequences were searched against the prokaryotic sequences in the GenEmbl database. The search identified 6 clones as part of the Omp2 gene, and 10 2 clones as part of the Omp3 gene, and 2 clones as part of the MOMP gene, indicating that COMC proteins had been successfully cloned. Furthermore, 32 clones were obtained, containing DNA sequences not found in the GenEmbl database. These sequences could, however, be clustered in two contigs 15 of 6 and 4 clones, and three clones were identical. In addition 19 clones were found with no overlap to the contigs (Figure 7). To obtain more sequence data for the genes, *C. pneumoniae* DNA was totally digested with BamHI restriction enzyme, and the fragments were cloned into the vector 20 pBluescript. The ligated DNA was electrotransformed into *E. coli* XL1-Blue and selected on plates containing Ampicillin. The recombinant bacterial colonies were transferred to a nitrocellulose membrane, and colony hybridisation was performed using the inserts of pEX 1-1 clone as a probe. A 25 clone containing a single BamHI fragment of 4.5 kb was found, and the hybridisation to the probe was confirmed by Southern blotting. The insert of the clone was sequenced bi-directionally using synthetic primers for approx. each 300 bp. The sequence of the BamHI fragment made it possible to 30 join the two contigs of pEX clones. Totally, together with the pEX clones it was possible to assemble 6.5 kb DNA sequence, encoding two new COMC proteins. (Figure 6)

Additional sequences were obtained by PCR performed on purified *C. pneumoniae* DNA with primers both from the known 35 Omp genes and from other known genes. The obtained PCR

products were sequenced, The sequence organisation is shown in Fig. 7. Additional 8 Omp genes were detected. The alignment of the deduced amino acid sequences are shown in Fig. 8 A and B.

## 5 Analysis of DNA sequence

The DNA sequence encoding the Omp4-15 proteins with a size of 89.6-100.3 kDa (and for Omp13: 56.1 kDa). Omp4 and Omp5 were transcribed in opposite directions. Downstream Omp4 a possible termination structure was located. The 3' end of the Omp5 gene was not cloned due to the presence of the BamHI restriction enzyme site positioned within the gene. The translated DNA sequence of Omp4 and Omp5 was compared by use of the gap programme in the GCG package (Wisconsin package, version 8.1-UNIX, August 1995, sequence analysis software package). The two genes had an amino acid identity of 41% (similarity 61%), and a possible cleavage site for signal peptidase 1 was present at amino acid 17 in Omp4 and amino acid 25 in Omp5. When the amino acid sequence encoded by two other pEX clones were compared to the sequence of Omp4 and Omp5 they also had amino acid homology to the genes. It is seen that the two clones have homology to the same area in the Omp4 and Omp5 proteins. Consequently, the pEX clones must have originated from two additional genes. Therefore these genes were named Omp6 and Omp7. Similar analyses were performed with the other genes. In contrast to what was seen for Omp4 and 5 none of the other putative omp proteins had a cleavage site for signal peptides.

### EXAMPLE 2

#### 30 Polyclonal monospecific antibodies against pEX fusion proteins and full length recombination + Omp4

To investigate the topology of the Omp4-7 proteins, representative pEX clones, were selected from each gene. The fusion proteins of  $\beta$ -galactosidase/omp were induced, and the

proteins were partially purified as inclusion bodies. Balb/c mice were immunized three times intramuscular with the antigens at an interval of one week, and after six weeks the serum was obtained from the mice. HeLa cells were infected with the *C. pneumoniae*. 72 hours after the infection the mono-layers were fixed with 3.7% formaldehyde. This treatment makes the outer membrane of the Chlamydia impermeable for antibodies due to the extensive cross-linking of the outer membrane proteins by the formaldehyde. The HeLa cells were permeabilized with 0.2% Triton X100, the monolayers were washed in PBS, then incubated with 20% (v/v) FCS to inactivate free radicals of the formaldehyde. The mice sera were diluted 1:100 PBS with 20% (v/v) FCS and incubated with the monolayers for half an hour. The monolayers were washed in PBS and secondary FITCH conjugated rabbit anti mouse serum was added for half an hour, and the monolayers were washed and mounted. Several of the antibodies reacted strongly with the EBs in the inclusions (Figure 9). In spite of the formaldehyde fixation it could not be excluded that the surface of the EB was changed by the treatments, so that the antibodies could get access to the Omp4-7. Therefore, the reaction was confirmed by immuno-electron microscopy with the antibody raised against clone pEX3-36. Purified EB of *C. pneumoniae* were absorbed to carbon coated nickel grids. After the absorption the grids were washed with PBS and blocked in 0.5% Ovalbumin dissolved in PBS. The antibodies were diluted 1:100 in the same buffer and incubated for 30 minutes. The grids were washed in PBS. Rabbit anti mouse Ig conjugated with 10nm colloidal gold diluted in PBS containing 1% gelatin was added to the grids for half an hour. The grids were washed in 3 x PBS with 1% gelatin and 3 times in PBS, the grids were contraststained with 0.7% phospho tungstic acid. The grids were analysed in a Jeol 1010 electron microscope at 40 kV. It was seen that the gold particles were covering the surface of the purified EB. Because the *C. pneumoniae* EBs were not exposed to any detergent or fixation under either the purification or the reaction with antibodies, these

results show that the cloned proteins have surface exposed epitopes.

#### Polyclonal monospecific antibodies against Omp4

The Omp4 gene was amplified by PCR with primers that  
5 contained LIC-sites, and the PCR product was cloned into the  
pET-30 LIC vector (Novagen). The histidine tagged fusion  
protein was expressed by induction of the synthesis by IPTG  
and purified over a nickel column. The purified Omp4 protein  
was used for immunization of a rabbit (six times, 8  $\mu$ g each  
10 time).

#### Use of rabbit polyclonal antibodies to recombinant Omp4 for detection of *Chlamydia pneumoniae* in paraffin embedded sections

The lungs of *C. pneumoniae* infected mice were obtained three  
15 days after intranasal infection. The tissue samples were  
fixed in 4% formaldehyde, paraffin embedded, sectioned and  
deparaffinized prior to staining. The sections were incubated  
with the rabbit serum diluted 1:200 in TBS ( 150 mM NaCl,  
20mM Tris pH 7.5) for 30 min at room temperature. After wash  
20 two times in TBS the sections were incubated with the  
secondary antibody (biotinylated goat anti-rabbit antibodies)  
diluted 1:300 in TBS, followed by two times wash in TBS. The  
sections were stained with streptavidin-biotin complex  
(streptABComplex/AP, Dako) for 30 min washed and developed  
25 under microscopic inspection with chromagen + new fuchsin  
(Vector laboratories). The sections were counter stained with  
Hematoxylin and analyzed ny microscopy.

#### Immuno blotting analysis with hyperimmune monospecific rabbit anti-serum

30 The insert of pEX1-1 clone was amplified by PCR using primers  
containing LIC sites. The PCR product could therefore be  
inserted in the pET-32 LIC vector (Novagen, UK cat No. 69076-



1). Thereby the insert sequence of the pEX1-1 clone was expressed in the new vector as a fusion protein, the part of the fusion protein encoded by the pET-32 LIC vector had 6 histidine residues in a row. The expression of the fusion protein was induced in this vector, and the fusion protein could be purified under denaturing condition on a Ni<sup>2+</sup> column due to the high affinity of the histidine residues to divalent cations. The purified protein was used for immunization of a New Zealand white rabbit. After 6 times intramuscular and 2 times intravenous immunization the serum was obtained from the rabbit. Purified *C. pneumoniae* EB was dissolved in SDS-sample buffer. Half of the sample was heated to 100°C in the sample buffer, whereas the other half of the sample was not heated. The samples were separated by SDS-PAGE, and the proteins were transferred to nitrocellulose, the serum was reacted with the strips. With the samples heated to 100°C the serum recognized a high molecular weight band of approximately 98 kDa. This is in agreement with the predicted size of Omp5, of which the pEX1-1 clone is a part, however, when the antibody was reacted to the strip with unheated EB, the pattern was different. Now a band was seen with a size of 75 kDa, in addition weaker bands were observed above the band (Figure 10). These data demonstrate that Omp5 needs boiling in SDS-sample buffer to be fully denatured and migrate with a size as predicted from the gene product. When the samples were not boiled, the protein was not fully denatured and less SDS binds to the protein and it has a more globular structure that will migrate faster in the acrylamide gel. The band pattern looked identical to what was obtained with a monoclonal antibody (MAb 26.1) (lane 6), we earlier have described (Christiansen et al., 1994), reacting with the surface of *C. pneumoniae* EB, but the antibody do not react with the fully SDS denatured *C. pneumoniae* EB in immunoblotting.

### Experimental infection of C57 black mice

Due to the realization of the altered migration of the Omp4-7 proteins without boiling, we chose to analyse antibodies against *C. pneumoniae* EBs after an experimental infection of mice. To obtain antibodies from an infection caused by *C. pneumoniae*, C57 black mice were inoculated intranasally with  $10^7$  CFI of *C. pneumoniae* under a light ether anaesthesia. After 14 days of infection the serum samples were obtained and the lungs were analysed for pathological changes. In two of the mice a severe pneumonia was observed in the lung sections, and in the third mouse only minor changes were observed. The serum from the mice was diluted-1:100 and reacted with purified EBs dissolved in sample buffer with and without boiling. In the preparations that had been heated to 100°C the sera from two of the mice reacted strongly with bands of 60/62 kDa and weaker bands of 55 kDa, but no reaction was observed with proteins of the size of Omp4-7 (Figure 11). However, when the sera were reacted with the preparation that had not been heated they all had a strong reaction with a broad band of an approximate size of 75 kDa. This is in agreement with the size of the Omp4-7 proteins in the unheated preparation. Therefore, it could be concluded that the epitopes of the Omp4-7 proteins recognized by the antibodies after a *C. pneumoniae* infection were discontinuous epitopes because the full denaturation of the antigen completely destroyed the epitopes. The 75 kDa protein observed in unheated samples is not Omp2 (Shown in immunoblotting with an Omp2 specific antibody)

### EXAMPLE 3

#### 30 Comparison of Omp4-7 of *C. pneumoniae* with putative outer membrane proteins (POMP) of *C. psittaci*

Longbottom et al. (1996) have published partial sequence from 98 to 90 kDa proteins from *C. psittaci*. They have entered the full sequence of 5 genes in this family in the EMBL database.

They have named the genes "putative outer membrane proteins" (POMP) since their precise location was not determined. The family is composed of two genes that are completely identical, and two genes with high homology to these genes.

5 They calculated a molecular size of 90 and 91 kDa. The 5th encode a protein of 98 kDa. The sequence of the Omp4-7 proteins of *C. pneumoniae* were compared to the sequences of the *C. Psittaci* POMP proteins with the programme pileup in the GCG package. The amino acid homologies were in the range  
10 of 51-63%. It is seen that the *C. pneumoniae* Omp4-5 proteins are most related to the 98 kDa POMP protein of *C. psittaci*. Interestingly, the 98 kDa *C. psittaci* POMP protein is more related to the *C. pneumoniae* genes than to the other *C. psittaci* genes. The repeated sequences of GGAI were conserved  
15 in the 98 kDa POMP protein, but only three GGAI repeats were present in the 90 and 91 kDa *C. psittaci* POMP proteins. For *C. psittaci* it has been shown that antibodies to these proteins seem to be protective for the infection.

#### REFERENCES

- 20
1. Caldwell, H.D., J. Kromhout and J. Schachter, Infect. Immun. 31, 1161-1176 (1981).
  2. Campbell, L.A., M.P. Melgosa, D.J. Hamilton, C.-C. Kuo and J.T. Grayston, J. Clinical Microbiol., 30,  
25 434-439 (1992).
  3. Christiansen, G., and S. Birkelund. Eur. Microbiol. 1:24-29 (1992).
  4. Christiansen, G., L. Østergaard, and S. Birkelund. Proceedings of the eight International symposium on  
30 Human Infections, Eds. Orfila et al., pp 173-176, (1994).
  5. Grayston, J.T., Kuo, C.-C., Campbell, L.A., and Vang, S.-P. Int. J. Syst. Bacteriol. 39, 88-90 (1989).
  6. Grayston, J.T., C.-C. Kuo, S.-P. Wang and J. Altman.  
35 1986. N. Engl. J. Med. 315, 161-168 (1986).
  7. Kuo, C.C., L.A. Jackson, L.A. Campbell and J.T. Graystone. Clin. Microbiol. Rev. 8, 451-461 (1995).

8. Longbottom, D., M. Russell, G.E Jones, A. Lainson, and A.J. Herring. FEMS Microbiol. Lett. 142, 277-281 (1996).
- 5 9. Melgosa, M.P., C.-C. Kuo and L.A. Campbell, FEMS Microbiol. Lett. 112, 199-204 (1993).
10. Campbell, L.A., C.-C kuo, S.P. Wang amd J.T. Grayston. J. Clin. Microbiol. 28, 1261-1264 (1990).
11. Halme, S., P. Saikku and H.-M. Surcel. Scand. J. Immunol. 45, 378-384 (1997).
- 10 12. Miyashita, N. and A. Matsumoto. J. Clin. Microbiol. 30, 2911-2916 (1992).
13. Wang, S.P., and J.T. Grayston, Am. J. Ophtalmol. 70, 367-374 (1970).
- 15 14. Freund, E.A., H. Ernø and R.M. Lemcke. Identification of mycoplasma, P377-443 in I. Norris and J.R. Bergen; Methods in Microbiology vol 13, A.P. Inc. London (1979)

## Claims (Amended)

1. Species specific diagnostic test for identifying infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said test comprising detecting in a patient or in a patient sample the presence of antibodies against one or more proteins from the outer membrane of *Chlamydia pneumoniae*, said proteins being outer membrane proteins selected from proteins having the sequence as shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or in SEQ ID NO: 24, or a variant or subsequence thereof or

being said proteins encoded by the nucleic acid fragments selected from nucleotide sequence shown in SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, or in SEQ ID NO: 23, or a variant or subsequence thereof and wherein a subsequence of 100 nucleic acids or lower shows a homology of at least 80% .

2. Diagnostic test according to claim 1 wherein detection of nucleic acid fragments is obtained by using nucleic acid amplification.

3. Diagnostic test according to claim 2, wherein detection of nucleic acid fragments is obtained by using polymerase chain reaction.

4. A nucleic acid fragment derived from *Chlamydia pneumoniae* comprising the nucleotide sequence SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, or SEQ ID NO: 23, or a variant or subsequence of said nucleotide sequence which has a sequence homology of at least 50% with any of the sequences mentioned and wherein a subsequence of 100 nucleic acids or lower shows a homology of at least 80% .

5. A protein derived from *Chlamydia pneumoniae* having the amino acid sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof having a sequence similarity of at least 50% and a similar biological function.

6. Polyclonal monospecific antibody against the protein with the sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12,

SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof.

7. A diagnostic kit for the diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said kit comprising a protein with the amino acid sequence SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof.

8. A diagnostic kit for the diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said kit comprising antibodies against a protein with the amino acid sequence SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof.

9. A diagnostic kit for the diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*, said kit comprising a nucleic acid fragment with the sequence SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, or SEQ ID NO: 23, or a variant or subsequence thereof and wherein a subsequence of 100 nucleic acids or lower shows a homology of at least 80% .

10. A composition for immunising a mammal, such as a human, against *Chlamydia pneumoniae*, said composition comprising a protein with the amino acid sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof.

11. Use of a protein with the sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof in diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*.

12. Use of the protein with the sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24 or a variant or

subsequence thereof in an undenatured form, in diagnosis of infection of a mammal, such as a human, with *Chlamydia pneumoniae*.

13. Use of a protein with the sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof, for immunising a mammal, such as a human, against *Chlamydia pneumoniae*.

14. Use of the protein with the sequence shown in SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6, SEQ ID NO: 8, SEQ ID NO: 10, SEQ ID NO: 12, SEQ ID NO: 14, SEQ ID NO: 16, SEQ ID NO: 18, SEQ ID NO: 20, SEQ ID NO: 22, or SEQ ID NO: 24, or a variant or subsequence thereof in an undenatured form, for immunising a mammal, such as a human, against *Chlamydia pneumoniae*.

15. Use of a nucleic acid fragment with the nucleotide sequence shown in SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5, SEQ ID NO: 7, SEQ ID NO: 9, SEQ ID NO: 11, SEQ ID NO: 13, SEQ ID NO: 15, SEQ ID NO: 17, SEQ ID NO: 19, SEQ ID NO: 21, or SEQ ID NO: 23, or a variant or subsequence of said nucleotide sequence which has a sequence homology of at least 50% and wherein a subsequence of 100 nucleic acids or lower shows a homology of at least 80% with any of the mentioned nucleotide sequences encoding a protein used for effecting *in vivo* expression of antigens against *Chlamydia pneumoniae*, in a mammal such as a human.

## ABSTRACT OF THE DISCLOSURE

The invention relates to the identification of members of a gene family from the human respiratory pathogen *Chlamydia pneumoniae*, encoding surface exposed membrane proteins of a size of approximately 89-101 kDa, preferably about 89.6-100.3 kDa and about 56.1 kDa. The invention relates to the novel DNA sequences, the deduced amino acid sequences of the corresponding proteins and the use of the DNA sequences and the proteins in diagnosis of infections caused by *C. pneumoniae*, in pathology, in epidemiology, and as vaccine components.



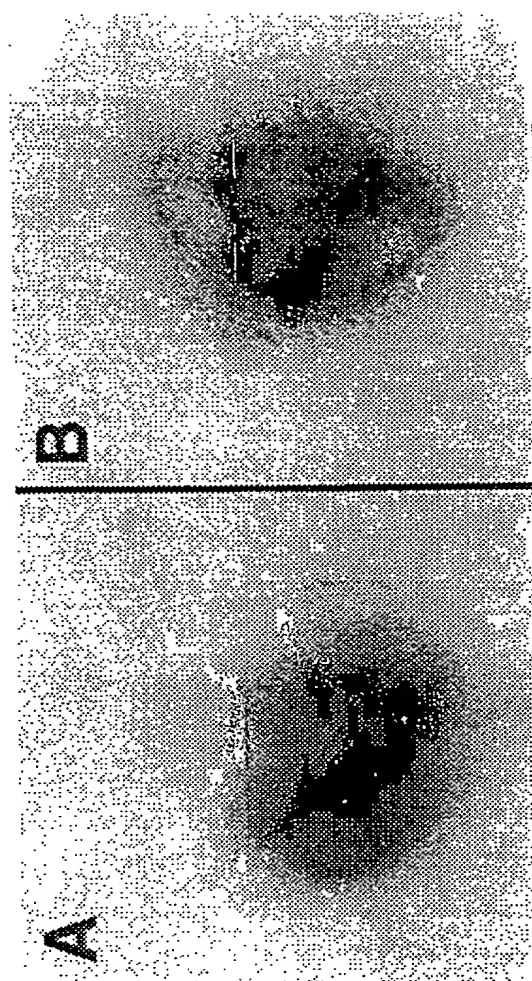


Fig. 1

2/21

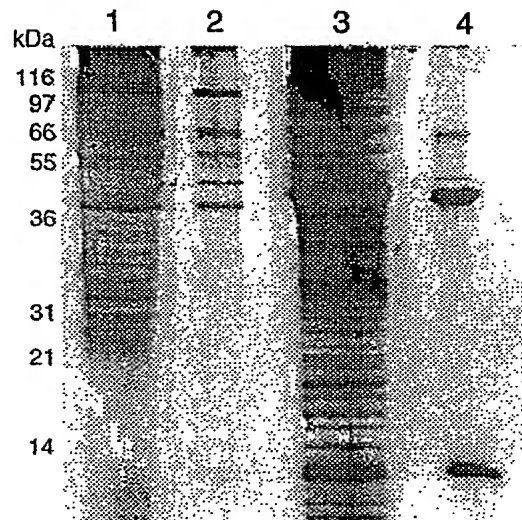


Fig. 2

3/21

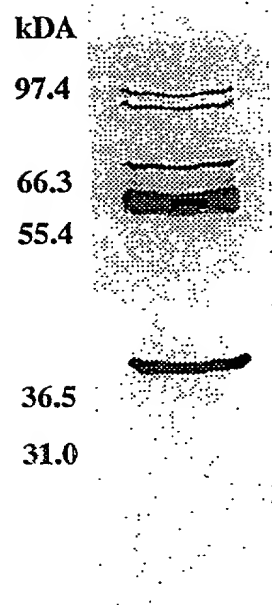


Fig. 3

4/21

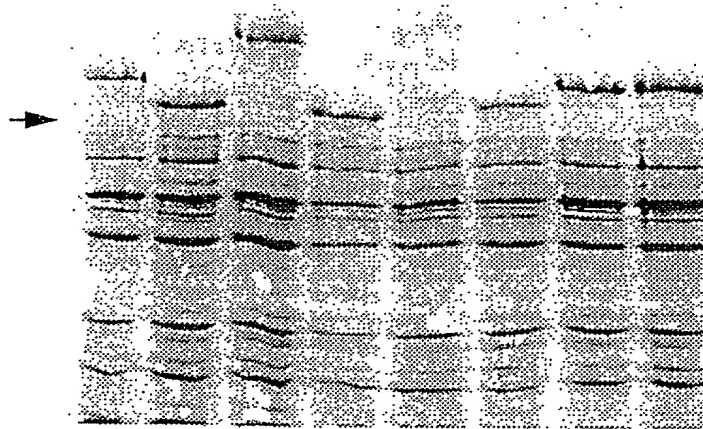


Fig. 4

5/21

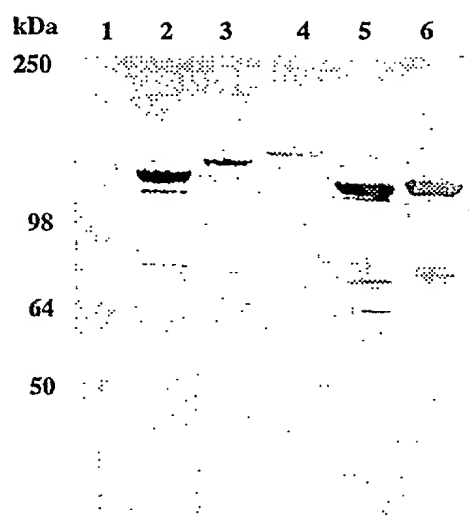


Fig. 5

6/21

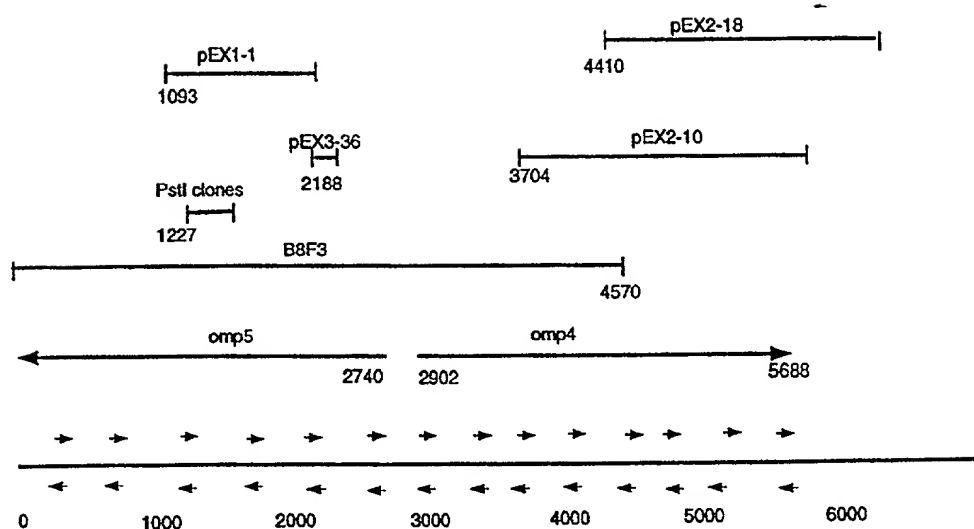


Fig. 6

7/21

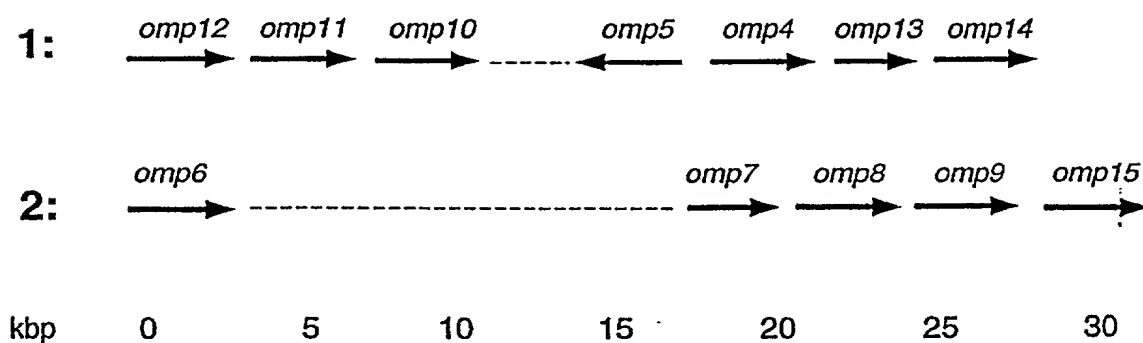
*C. pneumoniae omp4-15 gene clusters*

Fig. 7

5

[illegible]

**Fig. 8A**







[illegible][illegible]

**Fig. 8D**

12/21

[illegible][illegible]

**Fig. 8E**

•

[illegible]

Fig. 8F

[illegible][illegible]

**Fig. 8G**

[illegible]

**Fig. 8H.**

[illegible]



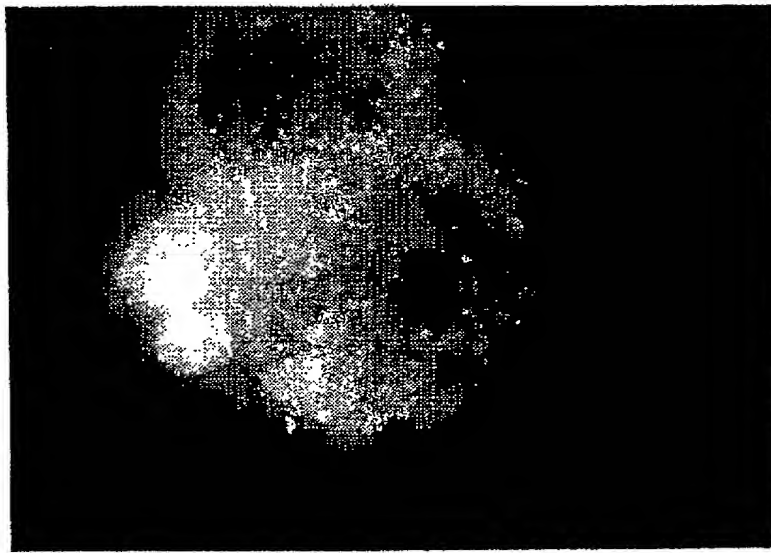
17/21

omp12 227 M K E S T K K S L - -  
 omp8 876 W W S W S W S W L - -  
 omp5 866 A S S W S W S W L - -  
 omp9 878 A S S W S W S W L - -  
 omp11 876 A S S W S W S W L - -  
 omp10 876 A S S W S W S W L - -  
 omp4 893 A S S W S W S W L - -  
 omp15 789 A S S W S W S W L - -  
 omp7 870 A S S W S W S W L - -  
 omp6 514 A S S W S W S W L - -  
 omp13 262 A S S W S W S W L - -  
 omp14 A S S W S W S W L - -  
 omp12 227 M K E S T K K S L - -  
 omp8 876 W W S W S W S W L - -  
 omp5 866 A S S W S W S W L - -  
 omp9 878 A S S W S W S W L - -  
 omp11 876 A S S W S W S W L - -  
 omp10 876 A S S W S W S W L - -  
 omp4 893 A S S W S W S W L - -  
 omp15 789 A S S W S W S W L - -  
 omp7 870 A S S W S W S W L - -  
 omp6 514 A S S W S W S W L - -  
 omp13 262 A S S W S W S W L - -  
 omp14 A S S W S W S W L - -  
 omp12 227 M K E S T K K S L - -  
 omp8 876 W W S W S W S W L - -  
 omp5 866 A S S W S W S W L - -  
 omp9 878 A S S W S W S W L - -  
 omp11 876 A S S W S W S W L - -  
 omp10 876 A S S W S W S W L - -  
 omp4 893 A S S W S W S W L - -  
 omp15 789 A S S W S W S W L - -  
 omp7 870 A S S W S W S W L - -  
 omp6 514 A S S W S W S W L - -  
 omp13 262 A S S W S W S W L - -  
 omp14 A S S W S W S W L - -

omp12 277 F Y F F Y F L - -  
 omp8 926 K K K R R K K I - -  
 omp5 926 S A G G L G G S S - -  
 omp9 916 G G G G L G G S S - -  
 omp11 928 T N D D L C L V L A G C - -  
 omp10 926 L L L L L L L L L L - -  
 omp4 926 T N V D D D D D D D D D - -  
 omp15 943 Y N Y N Y N Y N Y N - -  
 omp7 839 N N N I I S S S S S S S - -  
 omp6 920 S S S S S S S S S S S - -  
 omp13 514 R R R R R R R R R R R - -  
 omp14 262 V L L L L L L L L L L - -  
 omp12 277 F Y F F Y F L - -  
 omp8 926 K K K R R K K I - -  
 omp5 926 S A G G L G G S S - -  
 omp9 916 G G G G L G G S S - -  
 omp11 928 T N D D L C L V L A G C - -  
 omp10 926 L L L L L L L L L L - -  
 omp4 926 T N V D D D D D D D D D - -  
 omp15 943 Y N Y N Y N Y N Y N - -  
 omp7 839 N N N I I S S S S S S S - -  
 omp6 920 S S S S S S S S S S S - -  
 omp13 514 R R R R R R R R R R R - -  
 omp14 262 V L L L L L L L L L L - -  
 omp12 277 F Y F F Y F L - -  
 omp8 926 K K K R R K K I - -  
 omp5 926 S A G G L G G S S - -  
 omp9 916 G G G G L G G S S - -  
 omp11 928 T N D D L C L V L A G C - -  
 omp10 926 L L L L L L L L L L - -  
 omp4 926 T N V D D D D D D D D D - -  
 omp15 943 Y N Y N Y N Y N Y N - -  
 omp7 839 N N N I I S S S S S S S - -  
 omp6 920 S S S S S S S S S S S - -  
 omp13 514 R R R R R R R R R R R - -  
 omp14 262 V L L L L L L L L L L - -

omp12 279 C F F F F F F - -  
 omp8 928 Q F F F F F F - -  
 omp5 928 Q F F F F F F - -  
 omp9 918 G F F F F F F - -  
 omp11 930 S F F F F F F - -  
 omp10 928 Q F F F F F F - -  
 omp4 945 R F F F F F F - -  
 omp15 841 R F F F F F F - -  
 omp7 922 K F F F F F F - -  
 omp6 514 R F F F F F F - -  
 omp13 262 - - - - - - - - -  
 omp14 - - - - - - - - -  
 omp12 279 C F F F F F F - -  
 omp8 928 Q F F F F F F - -  
 omp5 928 Q F F F F F F - -  
 omp9 918 G F F F F F F - -  
 omp11 930 S F F F F F F - -  
 omp10 928 Q F F F F F F - -  
 omp4 945 R F F F F F F - -  
 omp15 841 R F F F F F F - -  
 omp7 922 K F F F F F F - -  
 omp6 514 R F F F F F F - -  
 omp13 262 - - - - - - - - -  
 omp14 - - - - - - - - -

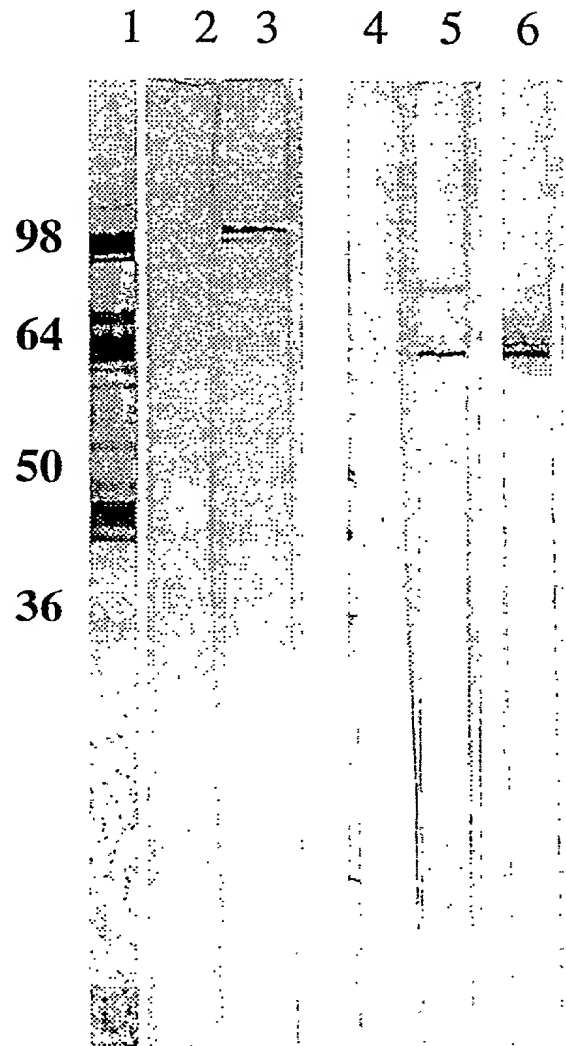
Fig. 8J



Omme sole var ikke  
 tilstede ved gennemgang  
 3/5-99.  
 Print fra Mimosa via  
 Dokumentations afd.

**Fig. 9**

19/21



Immunoblotting of *C. pneumoniae* EB, lane 1-3 heated to 100°C in SDS-sample buffer, lane 4-6 unheated. Lane 1 reacted with rabbit anti *C. pneumoniae* OMC; lane 2 and 4 pre-serum; lane 3 and 5 polyclonal rabbit anti pEX1-1 fusion protein; lane 6 MAb 26.1.

**Fig. 10**

20/21

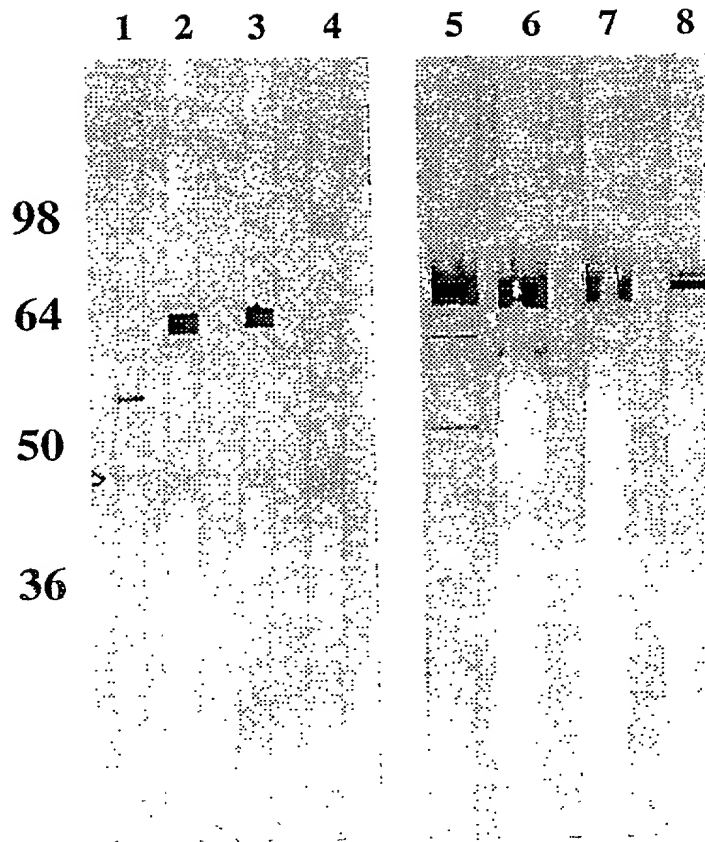


Fig. 11

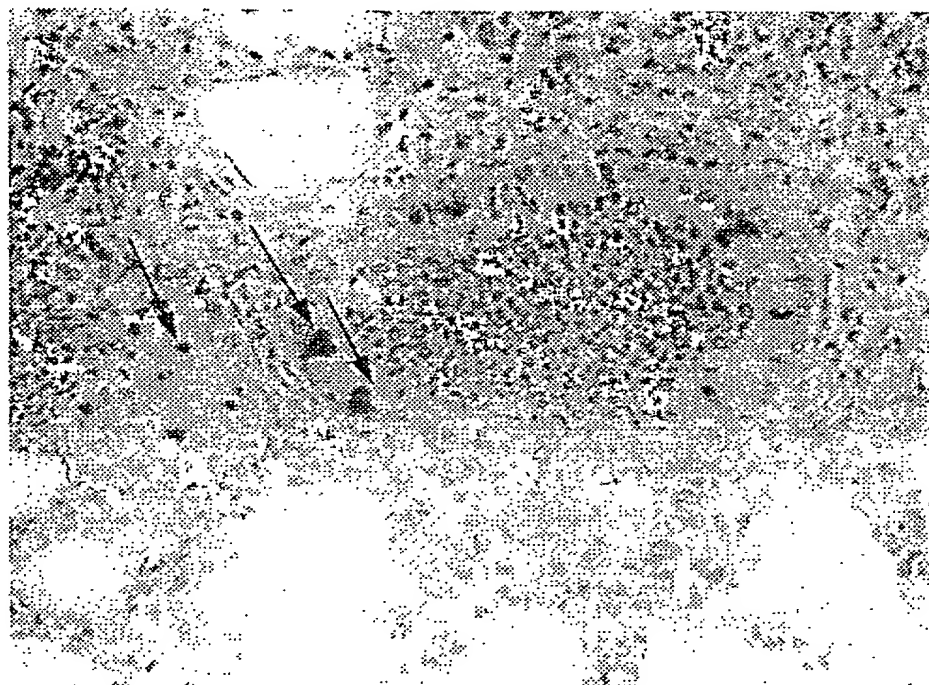


Fig. 12

**Combined Declaration for Patent Application and Power of Attorney**

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name; and that I believe I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled

Surface exposed proteins from Chlamydia Pneumoniae

\_\_\_\_\_ the specification of which (check one)

☐ is attached hereto;

☐ was filed in the United States under 35 U.S.C. §111 on \_\_\_\_\_, as  
USSN \_\_\_\_\_\*; or

☒ was/will be filed in the U.S. under 35 U.S.C. §371 by entry into the U.S. national stage of an international (PCT) application, PCT/DK98/00266; filed 19 June 1998, entry requested on \_\_\_\_\_\*; national stage application received  
USSN \_\_\_\_\_\*; §371/§102(e) date \_\_\_\_\_\* (\*if known),

and was amended on \_\_\_\_\_ (if applicable).

(include dates of amendments under PCT Art. 19 and 34 if PCT)

I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment referred to above; and I acknowledge the duty to disclose to the Patent and Trademark Office (PTO) all information known by me to be material to patentability as defined in 37 C.F.R. § 1.56.

I hereby claim foreign priority benefits under 35 U.S.C. §§ 119, 365 of any prior foreign application(s) for patent or inventor's certificate, or prior PCT application(s) designating a country other than the U.S., listed below with the "Yes" box checked and have also identified below any such application having a filing date before that of the application on which priority is claimed:

<u>0744/97</u>	<u>Denmark</u>	<u>23 June 1997</u>	<input checked="" type="checkbox"/>	<input type="checkbox"/>
(Number)	(Country)	(Day Month Year Filed)	YES	NO
_____	_____	_____	<input type="checkbox"/>	<input type="checkbox"/>
(Number)	(Country)	(Day Month Year Filed)	YES	NO

I hereby claim the benefit under 35 U.S.C. § 120 of any prior U.S. non-provisional Application(s) or prior PCT Application(s) designating the U.S. listed below, or under § 119(e) of any prior U.S. provisional applications listed below, and, insofar as the subject matter of each of the claims of this application is not disclosed in such U.S. or PCT application in the manner provided by the first paragraph of 35 U.S.C. §112, I acknowledge the duty to disclose to the PTO all information as defined in 37 C.F.R. §1.56(a) which occurred between the filing date of the prior application and the national filing date of this application:

_____ (Application Serial No.)	_____ (Day Month Year Filed)	_____ (Status: patented, pending, abandoned)
_____ (Application Serial No.)	_____ (Day Month Year Filed)	_____ (Status: patented, pending, abandoned)
_____ (Application Serial No.)	_____ (Day Month Year Filed)	_____ (Status: patented, pending, abandoned)

I hereby appoint the following attorneys, with full power of substitution, association, and revocation, to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith.

SHERIDAN NEIMARK, REG. NO. 20,520 - ROGER L. BROWDY, REG. NO. 25,618 - ANNE M. KORNBAU, REG. NO. 25,884  
NORMAN J. LATKER, REG. NO. 19,963 - IVER P. COOPER, REG. NO. 28,005 - ALLEN C. YUN, REG. NO. 37,971\* 7  
NICK S. BROMER, REG. NO. 33,478 - \* Patent Agent

ADDRESS ALL CORRESPONDENCE TO  
**BROWDY AND NEIMARK, P.L.L.C.**  
419 Seventh Street, N.W.  
Washington, D.C. 20004

DIRECT ALL TELEPHONE CALLS TO:  
**BROWDY AND NEIMARK**  
**(202) 628-5197**

The undersigned hereby authorizes the U.S. Attorneys or Agents named herein to accept and follow instructions from **PLOUGMANN, VINGTOFT & PARTNERS** as to any action to be taken in the U.S. Patent and Trademark Office regarding this application without direct communication between the U.S. Attorney or Agent and the undersigned. In the event of a change of the persons from whom instructions may be taken, the U.S. Attorneys or Agents named herein will be so notified by the undersigned.

Title: Surface exposed proteins from Chlamydia Pneumoniae

U.S. Application filed \_\_\_\_\_, Serial No. \_\_\_\_\_

PCT Application filed 19 June 1998, Serial No. PCT/DK98/00266

I hereby further declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under 18 U.S.C. §1001 and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

FULL NAME OF FIRST INVENTOR <u>Svend Birkelund</u>		INVENTOR'S SIGNATURE <u>Svend Birkelund</u>	DATE <u>10/3-2000</u>
RESIDENCE <u>Egå, Denmark</u> <u>DKX</u>		CITIZENSHIP Danish	
POST OFFICE ADDRESS <u>Søtoften 26, DK-8250 Egå, Denmark</u>			
FULL NAME OF SECOND JOINT INVENTOR <u>Gunna Christiansen</u>		INVENTOR'S SIGNATURE <u>Gunna Christensen</u>	DATE <u>10/3-2000</u>
RESIDENCE <u>Egå, Denmark</u> <u>DKX</u>		CITIZENSHIP Danish	
POST OFFICE ADDRESS <u>Søtoften 26, DK-8250 Egå, Denmark</u>			
FULL NAME OF THIRD JOINT INVENTOR <u>Anna-Sofie Hebsgaard Pedersen</u>		INVENTOR'S SIGNATURE <u>Anna-Sofie H Pedersen</u>	DATE <u>10/3-00</u>
RESIDENCE <u>Silkeborg, Denmark</u> <u>DKX</u>		CITIZENSHIP Danish	
POST OFFICE ADDRESS <u>Vestergade 26C, 2.th., DK-8600 Silkeborg, Denmark</u>			
FULL NAME OF FOURTH JOINT INVENTOR <u>Per Mygind</u>		INVENTOR'S SIGNATURE <u>Per Mygind</u>	DATE <u>10/3-2000</u>
RESIDENCE <u>Århus, Denmark</u> <u>DKX</u>		CITIZENSHIP Danish	
POST OFFICE ADDRESS <u>Cort Adelers Gade 17, 1. tv., DK-8200 Århus N, Denmark</u>			
FULL NAME OF FIFTH JOINT INVENTOR <u>Katrine Knudsen</u>		INVENTOR'S SIGNATURE <u>Katrine Knudsen</u>	DATE <u>10-3-2000</u>
RESIDENCE <u>Århus, Denmark</u> <u>DKX</u>		CITIZENSHIP Danish	
POST OFFICE ADDRESS <u>Lundingsgade 33, lejlighed 407, DK-8000 Århus C, Denmark</u>			
FULL NAME OF SIXTH JOINT INVENTOR		INVENTOR'S SIGNATURE	DATE
RESIDENCE		CITIZENSHIP	
POST OFFICE ADDRESS			
FULL NAME OF SEVENTH JOINT INVENTOR		INVENTOR'S SIGNATURE	DATE
RESIDENCE		CITIZENSHIP	
POST OFFICE ADDRESS			

## SEQUENCE LISTING

## (1) GENERAL INFORMATION

## (i) APPLICANT

- (A) NAME: Svend Birkelund  
 (B) STREET: Dept. of Medical Microbiology and Immunology,  
 University of Århus  
 (C) CITY: Århus C  
 (D) STATE OR PROVINCE:  
 (E) COUNTRY: Denmark  
 (F) POSTAL CODE: 8000

(ii) TITLE OF THE INVENTION: Chlamydia pneumoniae anti  
 gens

(iii) NUMBER OF SEQUENCES: 30

## (iv) COMPUTER-READABLE FORM:

- (A) MEDIUM TYPE: Diskette  
 (B) COMPUTER: IBM Compatible  
 (C) OPERATING SYSTEM: DOS  
 (D) SOFTWARE: FastSEQ for Windows Version 2.0

## (v) CURRENT APPLICATION DATA:

(A) APPLICATION NUMBER:

## (2) INFORMATION FOR SEQ ID NO:1:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 3200 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ix) FEATURE:

- (A) NAME/KEY: Coding Sequence  
 (B) LOCATION: 205...2987  
 (D) OTHER INFORMATION:

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

CAATGTCGAA	GAGAGCACTA	ACCAGGAAAA	TTGCGATTTC	ATAAACCCAC	TTTATTATTA	60
AATTCTTACT	TGCGTCATAT	AAAATAGAAA	ACTCAGAGAG	TCAAGATAAA	AATTCTTGAC	120
AGCTGTTTTG	TCATCTTTAA	CTTGATTAC	TTATTTGTT	TCTATATTGA	TGCGAATAGT	180
TCTCTAAAAA	ACAAAAGCAT	TACC ATG AAG	ACT TCG ATT	CCT TGG GTT	TTA	231
		Met Lys Thr Ser Ile Pro Trp Val Leu				
		1	5			
GTT TCC TCC	GTG TTA GCT	TTC TCA TGT	CAC CTA CAG	TCA CTA GCT	AAC	279
Val Ser Ser	Val Leu Ala	Phe Ser Cys	His Leu Gln	Ser Leu Ala	Asn	
10	15	20	25			



GAG GAA CTT TTA TCA CCT GAT GAT AGC TTT AAT GGA AAT ATC GAT TCA	327
Glu Glu Leu Leu Ser Pro Asp Asp Ser Phe Asn Gly Asn Ile Asp Ser	
30 35 40	
GGA ACG TTT ACT CCA AAA ACT TCA GCC ACA ACA TAT TCT CTA ACA GGA	375
Gly Thr Phe Thr Pro Lys Thr Ser Ala Thr Thr Tyr Ser Leu Thr Gly	
45 50 55	
GAT GTC TTC TTT TAC GAG CCT GGA AAA GGC ACT CCC TTA TCT GAC AGT	423
Asp Val Phe Phe Tyr Glu Pro Gly Lys Gly Thr Pro Leu Ser Asp Ser	
60 65 70	
TGT TTT AAG CAA ACC ACG GAC AAT CTT ACC TTC TTG GGG AAC GGT CAT	471
Cys Phe Lys Gln Thr Thr Asp Asn Leu Thr Phe Leu Gly Asn Gly His	
75 80 85	
AGC TTA ACG TTT GGC TTT ATA GAT GCT GGC ACT CAT GCA GGT GCT GCT	519
Ser Leu Thr Phe Gly Phe Ile Asp Ala Gly Thr His Ala Gly Ala Ala	
90 95 100 105	
GCA TCT ACA ACA GCA AAT AAG AAT CTT ACC TTC TCA GGG TTT TCC TTA	567
Ala Ser Thr Thr Ala Asn Lys Asn Leu Thr Phe Ser Gly Phe Ser Leu	
110 115 120	
CTG AGT TTT GAT TCC TCT CCT AGC ACA ACG GTT ACT ACA GGT CAG GGA	615
Leu Ser Phe Asp Ser Ser Pro Ser Thr Thr Val Thr Thr Gly Gln Gly	
125 130 135	
ACG CTT TCC TCA GCA GGA GGC GTA AAT TTA GAA AAT ATT CGT AAA CTT	663
Thr Leu Ser Ser Ala Gly Gly Val Asn Leu Glu Asn Ile Arg Lys Leu	
140 145 150	
GTA GTT GCT GGG AAT TTT TCT ACT GCA GAT GGT GGA GCT ATC AAA GGA	711
Val Val Ala Gly Asn Phe Ser Thr Ala Asp Gly Gly Ala Ile Lys Gly	
155 160 165	
GCG TCT TTC CTT TTA ACT GGC ACT TCT GGA GAT GCT CTT TTT AGT AAC	759
Ala Ser Phe Leu Leu Thr Gly Thr Ser Gly Asp Ala Leu Phe Ser Asn	
170 175 180 185	
AAC TCT TCA TCA ACA AAG GGA GGA GCA ATT GCT ACT ACA GCA GGC GCT	807
Asn Ser Ser Ser Thr Lys Gly Gly Ala Ile Ala Thr Thr Ala Gly Ala	
190 195 200	
CGC ATA GCA AAT AAC ACA GGT TAT GTT AGA TTC CTA TCT AAC ATA GCG	855
Arg Ile Ala Asn Asn Thr Gly Tyr Val Arg Phe Leu Ser Asn Ile Ala	
205 210 215	
TCT ACG TCA GGA GGC GCT ATC GAT GAT GAA GGC ACG TCG ATA CTA TCG	903
Ser Thr Ser Gly Gly Ala Ile Asp Asp Glu Gly Thr Ser Ile Leu Ser	
220 225 230	
AAC AAC AAA TTT CTA TAT TTT GAA GGG AAT GCA GCG AAA ACT ACT GGC	951
Asn Asn Lys Phe Leu Tyr Phe Glu Gly Asn Ala Ala Lys Thr Thr Gly	
235 240 245	
GGT GCG ATC TGC AAC ACC AAG GCG AGT GGA TCT CCT GAA CTG ATA ATC	999

Gly	Ala	Ile	Cys	Asn	Thr	Lys	Ala	Ser	Gly	Ser	Pro	Glu	Leu	Ile	Ile		
250					255				260						265		
TCT	AAC	AAT	AAG	ACT	CTG	ATC	TTT	GCT	TCA	AAC	GTA	GCA	GAA	ACA	AGC	1047	
Ser	Asn	Asn	Lys	Thr	Leu	Ile	Phe	Ala	Ser	Asn	Val	Ala	Glu	Thr	Ser		
				270				275						280			
GGT	GGC	GCC	ATC	CAT	GCT	AAA	AAG	CTA	GCC	CTT	TCC	TCT	GGA	GGC	TTT	1095	
Gly	Gly	Ala	Ile	His	Ala	Lys	Lys	Leu	Ala	Leu	Ser	Ser	Gly	Gly	Phe		
			285					290					295				
ACA	GAG	TTT	CTA	CGA	AAT	AAT	GTC	TCA	TCA	GCA	ACT	CCT	AAG	GGG	GGT	1143	
Thr	Glu	Phe	Leu	Arg	Asn	Asn	Val	Ser	Ser	Ala	Thr	Pro	Lys	Gly	Gly		
		300					305					310					
GCT	ATC	AGC	ATC	GAT	GCC	TCA	GGA	GAG	CTC	AGT	CTT	TCT	GCA	GAG	ACA	1191	
Ala	Ile	Ser	Ile	Asp	Ala	Ser	Gly	Glu	Leu	Ser	Leu	Ser	Ala	Glu	Thr		
	315					320					325						
GGA	AAC	ATT	ACC	TTT	GTA	AGA	AAT	ACC	CTT	ACA	ACA	ACC	GGA	AGT	ACC	1239	
Gly	Asn	Ile	Thr	Phe	Val	Arg	Asn	Thr	Leu	Thr	Thr	Thr	Gly	Ser	Thr		
330					335				340					345			
GAT	ACT	CCT	AAA	CGT	AAT	GCG	ATC	AAC	ATA	GGA	AGT	AAC	GGG	AAA	TTC	1287	
Asp	Thr	Pro	Lys	Arg	Asn	Ala	Ile	Asn	Ile	Gly	Ser	Asn	Gly	Lys	Phe		
				350				355						360			
ACG	GAA	TTA	CGG	GCT	GCT	AAA	AAT	CAT	ACA	ATT	TTC	TTC	TAT	GAT	CCC	1335	
Thr	Glu	Leu	Arg	Ala	Ala	Lys	Asn	His	Thr	Ile	Phe	Phe	Tyr	Asp	Pro		
			365					370					375				
ATC	ACT	TCA	GAA	GGA	ACC	TCA	TCA	GAC	GTA	TTG	AAG	ATA	AAT	AAC	GGC	1383	
Ile	Thr	Ser	Glu	Gly	Thr	Ser	Ser	Asp	Val	Leu	Lys	Ile	Asn	Asn	Gly		
		380					385					390					
TCT	GCG	GGA	GCT	CTC	AAT	CCA	TAT	CAA	GGA	ACG	ATT	CTA	TTT	TCT	GGA	1431	
Ser	Ala	Gly	Ala	Leu	Asn	Pro	Tyr	Gln	Gly	Thr	Ile	Leu	Phe	Ser	Gly		
	395					400					405						
GAA	ACC	CTA	ACA	GCA	GAT	GAA	CTT	AAA	GTT	GCT	GAC	AAT	TTA	AAA	TCT	1479	
Glu	Thr	Leu	Thr	Ala	Asp	Glu	Leu	Lys	Val	Ala	Asp	Asn	Leu	Lys	Ser		
410					415					420				425			
TCA	TTC	ACG	CAG	CCA	GTC	TCC	CTA	TCC	GGA	GGA	AAG	TTA	TTG	CTA	CAA	1527	
Ser	Phe	Thr	Gln	Pro	Val	Ser	Leu	Ser	Gly	Gly	Lys	Leu	Leu	Leu	Gln		
				430					435					440			
AAG	GGA	GTC	ACT	TTA	GAG	AGC	ACG	AGC	TTC	TCT	CAA	GAG	GCC	GGT	TCT	1575	
Lys	Gly	Val	Thr	Leu	Glu	Ser	Thr	Ser	Phe	Ser	Gln	Glu	Ala	Gly	Ser		
			445					450					455				
CTC	CTC	GGC	ATG	GAT	TCA	GGA	ACG	ACA	TTA	TCA	ACT	ACA	GCT	GGG	AGT	1623	
Leu	Leu	Gly	Met	Asp	Ser	Gly	Thr	Thr	Leu	Ser	Thr	Thr	Ala	Gly	Ser		
		460					465					470					
ATT	ACA	ATC	ACG	AAC	CTA	GGA	ATC	AAT	GTT	GAC	TCC	TTA	GGT	CTT	AAG	1671	
Ile	Thr	Ile	Thr	Asn	Leu	Gly	Ile	Asn	Val	Asp	Ser	Leu	Gly	Leu	Lys		

475	480	485	
CAG CCC GTC AGC CTA ACA GCA AAA GGT GCT TCA AAT AAA GTG ATC GTA Gln Pro Val Ser Leu Thr Ala Lys Gly Ala Ser Asn Lys Val Ile Val 490 495 500 505	1719		
TCT GGG AAG CTC AAC CTG ATT GAT ATT GAA GGG AAC ATT TAT GAA AGT Ser Gly Lys Leu Asn Leu Ile Asp Ile Glu Gly Asn Ile Tyr Glu Ser 510 515 520	1767		
CAT ATG TTC AGC CAT GAC CAG CTC TTC TCT CTA TTA AAA ATC ACG GTT His Met Phe Ser His Asp Gln Leu Phe Ser Leu Leu Lys Ile Thr Val 525 530 535	1815		
GAT GCT GAT GTT GAT ACT AAC GTT GAC ATC AGC AGC CTT ATC CCT GTT Asp Ala Asp Val Asp Thr Asn Val Asp Ile Ser Ser Leu Ile Pro Val 540 545 550	1863		
CCT GCT GAG GAT CCT AAT TCA GAA TAC GGA TTC CAA GGA CAA TGG AAT Pro Ala Glu Asp Pro Asn Ser Glu Tyr Gly Phe Gln Gly Gln Trp Asn 555 560 565	1911		
GTT AAT TGG ACT ACG GAT ACA GCT ACA AAT ACA AAA GAG GCC ACG GCA Val Asn Trp Thr Thr Asp Thr Ala Thr Asn Thr Lys Glu Ala Thr Ala 570 575 580 585	1959		
ACT TGG ACC AAA ACA GGA TTT GTT CCC AGC CCC GAA AGA AAA TCT GCG Thr Trp Thr Lys Thr Gly Phe Val Pro Ser Pro Glu Arg Lys Ser Ala 590 595 600	2007		
TTA GTA TGC AAT ACC CTA TGG GGA GTC TTT ACT GAC ATT CGC TCT CTG Leu Val Cys Asn Thr Leu Trp Gly Val Phe Thr Asp Ile Arg Ser Leu 605 610 615	2055		
CAA CAG CTT GTA GAG ATC GGC GCA ACT GGT ATG GAA CAC AAA CAA GGT Gln Gln Leu Val Glu Ile Gly Ala Thr Gly Met Glu His Lys Gln Gly 620 625 630	2103		
TTC TGG GTT TCC TCC ATG ACG AAC TTC CTG CAT AAG ACT GGA GAT GAA Phe Trp Val Ser Ser Met Thr Asn Phe Leu His Lys Thr Gly Asp Glu 635 640 645	2151		
AAT CGC AAA GGC TTC CGT CAT ACC TCT GGA GGC TAC GTC ATC GGT GGA Asn Arg Lys Gly Phe Arg His Thr Ser Gly Gly Tyr Val Ile Gly Gly 650 655 660 665	2199		
AGT GCT CAC ACT CCT AAA GAC GAC CTA TTT ACC TTT GCG TTC TGC CAT Ser Ala His Thr Pro Lys Asp Asp Leu Phe Thr Phe Ala Phe Cys His 670 675 680	2247		
CTC TTT GCT AGA GAC AAA GAT TGT TTT ATC GCT CAC AAC AAC TCT AGA Leu Phe Ala Arg Asp Lys Asp Cys Phe Ile Ala His Asn Asn Ser Arg 685 690 695	2295		
ACC TAC GGT GGA ACT TTA TTC TTC AAG CAC TCT CAT ACC CTA CAA CCC Thr Tyr Gly Gly Thr Leu Phe Phe Lys His Ser His Thr Leu Gln Pro 700 705 710	2343		

CAA AAC TAT TTG AGA TTA GGA AGA GCA AAG TTT TCT GAA TCA GCT ATA 2391  
 Gln Asn Tyr Leu Arg Leu Gly Arg Ala Lys Phe Ser Glu Ser Ala Ile  
 715 720 725

GAA AAA TTC CCT AGG GAA ATT CCC CTA GCC TTG GAT GTC CAA GTT TCG 2439  
 Glu Lys Phe Pro Arg Glu Ile Pro Leu Ala Leu Asp Val Gln Val Ser  
 730 735 740 745

TTC AGC CAT TCA GAC AAC CGT ATG GAA ACG CAC TAT ACC TCA TTG CCA 2487  
 Phe Ser His Ser Asp Asn Arg Met Glu Thr His Tyr Thr Ser Leu Pro  
 750 755 760

GAA TCC GAA GGT TCT TGG AGC AAC GAG TGT ATA GCT GGT GGT ATC GGC 2535  
 Glu Ser Glu Gly Ser Trp Ser Asn Glu Cys Ile Ala Gly Gly Ile Gly  
 765 770 775

CTA GAC CTT CCT TTT GTT CTT TCC AAC CCA CAT CCT CTT TTC AAG ACC 2583  
 Leu Asp Leu Pro Phe Val Leu Ser Asn Pro His Pro Leu Phe Lys Thr  
 780 785 790

TTC ATT CCA CAG ATG AAA GTC GAA ATG GTT TAT GTA TCA CAA AAT AGC 2631  
 Phe Ile Pro Gln Met Lys Val Glu Met Val Tyr Val Ser Gln Asn Ser  
 795 800 805

TTC TTC GAA AGC TCT AGT GAT GGC CGT GGT TTT AGT ATT GGA AGG CTG 2679  
 Phe Phe Glu Ser Ser Ser Asp Gly Arg Gly Phe Ser Ile Gly Arg Leu  
 810 815 820 825

CTT AAC CTC TCG ATT CCT GTG GGT GCG AAA TTC GTG CAG GGG GAT ATC 2727  
 Leu Asn Leu Ser Ile Pro Val Gly Ala Lys Phe Val Gln Gly Asp Ile  
 830 835 840

GGA GAT TCC TAC ACC TAT GAT CTC TCA GGA TTC TTT GTT TCC GAT GTC 2775  
 Gly Asp Ser Tyr Thr Tyr Asp Leu Ser Gly Phe Phe Val Ser Asp Val  
 845 850 855

TAT CGT AAC AAT CCC CAA TCT ACA GCG ACT CTT GTG ATG AGC CCA GAC 2823  
 Tyr Arg Asn Asn Pro Gln Ser Thr Ala Thr Leu Val Met Ser Pro Asp  
 860 865 870

TCT TGG AAA ATT CGC GGT GGC AAT CTT TCA AGA CAG GCA TTT TTA CTG 2871  
 Ser Trp Lys Ile Arg Gly Gly Asn Leu Ser Arg Gln Ala Phe Leu Leu  
 875 880 885

AGG GGT AGC AAC AAC TAC GTC TAC AAC TCC AAT TGT GAG CTC TTC GGA 2919  
 Arg Gly Ser Asn Asn Tyr Val Tyr Asn Ser Asn Cys Glu Leu Phe Gly  
 890 895 900 905

CAT TAC GCT ATG GAA CTC CGT GGA TCT TCA AGG AAC TAC AAT GTA GAT 2967  
 His Tyr Ala Met Glu Leu Arg Gly Ser Ser Arg Asn Tyr Asn Val Asp  
 910 915 920

GTT GGT ACC AAA CTC CGA TT CTAGATTGCT AAAACTCCCT AGTTCTTCTA GGGAG 3022  
 Val Gly Thr Lys Leu Arg Phe  
 925

TTTTCTCATA CTTTtaggga AATATTGCT ATAGGGAATG CTTTCCTTGC AAAGTGTAAA 3082

AAATAACATT TGTCCCTCTT CAAAAAAGAT TTCTTTTAAT AATTTCTAGT TATAATTTTA 3142  
 TTTTAAAAAC AGTTAAATAA TTAATAGACA ATAATCTATT CTTATTGACT TCTTTTTT 3200

## (2) INFORMATION FOR SEQ ID NO:2:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 928 amino acids  
 (B) TYPE: amino acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: protein

## (v) FRAGMENT TYPE: internal

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

Met Lys Thr Ser Ile Pro Trp Val Leu Val Ser Ser Val Leu Ala Phe  
 1 5 10 15  
 Ser Cys His Leu Gln Ser Leu Ala Asn Glu Glu Leu Leu Ser Pro Asp  
 20 25 30  
 Asp Ser Phe Asn Gly Asn Ile Asp Ser Gly Thr Phe Thr Pro Lys Thr  
 35 40 45  
 Ser Ala Thr Thr Tyr Ser Leu Thr Gly Asp Val Phe Phe Tyr Glu Pro  
 50 55 60  
 Gly Lys Gly Thr Pro Leu Ser Asp Ser Cys Phe Lys Gln Thr Thr Asp  
 65 70 75 80  
 Asn Leu Thr Phe Leu Gly Asn Gly His Ser Leu Thr Phe Gly Phe Ile  
 85 90 95  
 Asp Ala Gly Thr His Ala Gly Ala Ala Ala Ser Thr Thr Ala Asn Lys  
 100 105 110  
 Asn Leu Thr Phe Ser Gly Phe Ser Leu Leu Ser Phe Asp Ser Ser Pro  
 115 120 125  
 Ser Thr Thr Val Thr Thr Gly Gln Gly Thr Leu Ser Ser Ala Gly Gly  
 130 135 140  
 Val Asn Leu Glu Asn Ile Arg Lys Leu Val Val Ala Gly Asn Phe Ser  
 145 150 155 160  
 Thr Ala Asp Gly Gly Ala Ile Lys Gly Ala Ser Phe Leu Leu Thr Gly  
 165 170 175  
 Thr Ser Gly Asp Ala Leu Phe Ser Asn Asn Ser Ser Ser Thr Lys Gly  
 180 185 190  
 Gly Ala Ile Ala Thr Thr Ala Gly Ala Arg Ile Ala Asn Asn Thr Gly  
 195 200 205  
 Tyr Val Arg Phe Leu Ser Asn Ile Ala Ser Thr Ser Gly Gly Ala Ile  
 210 215 220  
 Asp Asp Glu Gly Thr Ser Ile Leu Ser Asn Asn Lys Phe Leu Tyr Phe  
 225 230 235 240  
 Glu Gly Asn Ala Ala Lys Thr Thr Gly Gly Ala Ile Cys Asn Thr Lys  
 245 250 255  
 Ala Ser Gly Ser Pro Glu Leu Ile Ile Ser Asn Asn Lys Thr Leu Ile  
 260 265 270  
 Phe Ala Ser Asn Val Ala Glu Thr Ser Gly Gly Ala Ile His Ala Lys  
 275 280 285  
 Lys Leu Ala Leu Ser Ser Gly Gly Phe Thr Glu Phe Leu Arg Asn Asn  
 290 295 300  
 Val Ser Ser Ala Thr Pro Lys Gly Gly Ala Ile Ser Ile Asp Ala Ser

305                      310                      315                      320  
 Gly Glu Leu Ser Leu Ser Ala Glu Thr Gly Asn Ile Thr Phe Val Arg  
                                  325                      330                      335  
 Asn Thr Leu Thr Thr Thr Gly Ser Thr Asp Thr Pro Lys Arg Asn Ala  
                                  340                      345                      350  
 Ile Asn Ile Gly Ser Asn Gly Lys Phe Thr Glu Leu Arg Ala Ala Lys  
                                  355                      360                      365  
 Asn His Thr Ile Phe Phe Tyr Asp Pro Ile Thr Ser Glu Gly Thr Ser  
                                  370                      375                      380  
 Ser Asp Val Leu Lys Ile Asn Asn Gly Ser Ala Gly Ala Leu Asn Pro  
 385                                   390                      395                      400  
 Tyr Gln Gly Thr Ile Leu Phe Ser Gly Glu Thr Leu Thr Ala Asp Glu  
                                  405                      410                      415  
 Leu Lys Val Ala Asp Asn Leu Lys Ser Ser Phe Thr Gln Pro Val Ser  
                                  420                      425                      430  
 Leu Ser Gly Gly Lys Leu Leu Leu Gln Lys Gly Val Thr Leu Glu Ser  
                                  435                      440                      445  
 Thr Ser Phe Ser Gln Glu Ala Gly Ser Leu Leu Gly Met Asp Ser Gly  
                                  450                      455                      460  
 Thr Thr Leu Ser Thr Thr Ala Gly Ser Ile Thr Ile Thr Asn Leu Gly  
 465                                   470                      475                      480  
 Ile Asn Val Asp Ser Leu Gly Leu Lys Gln Pro Val Ser Leu Thr Ala  
                                  485                      490                      495  
 Lys Gly Ala Ser Asn Lys Val Ile Val Ser Gly Lys Leu Asn Leu Ile  
                                  500                      505                      510  
 Asp Ile Glu Gly Asn Ile Tyr Glu Ser His Met Phe Ser His Asp Gln  
                                  515                      520                      525  
 Leu Phe Ser Leu Leu Lys Ile Thr Val Asp Ala Asp Val Asp Thr Asn  
                                  530                      535                      540  
 Val Asp Ile Ser Ser Leu Ile Pro Val Pro Ala Glu Asp Pro Asn Ser  
 545                                   550                      555                      560  
 Glu Tyr Gly Phe Gln Gly Gln Trp Asn Val Asn Trp Thr Thr Asp Thr  
                                  565                      570                      575  
 Ala Thr Asn Thr Lys Glu Ala Thr Ala Thr Trp Thr Lys Thr Gly Phe  
                                  580                      585                      590  
 Val Pro Ser Pro Glu Arg Lys Ser Ala Leu Val Cys Asn Thr Leu Trp  
                                  595                      600                      605  
 Gly Val Phe Thr Asp Ile Arg Ser Leu Gln Gln Leu Val Glu Ile Gly  
                                  610                      615                      620  
 Ala Thr Gly Met Glu His Lys Gln Gly Phe Trp Val Ser Ser Met Thr  
 625                                   630                      635                      640  
 Asn Phe Leu His Lys Thr Gly Asp Glu Asn Arg Lys Gly Phe Arg His  
                                  645                      650                      655  
 Thr Ser Gly Gly Tyr Val Ile Gly Gly Ser Ala His Thr Pro Lys Asp  
                                  660                      665                      670  
 Asp Leu Phe Thr Phe Ala Phe Cys His Leu Phe Ala Arg Asp Lys Asp  
                                  675                      680                      685  
 Cys Phe Ile Ala His Asn Asn Ser Arg Thr Tyr Gly Gly Thr Leu Phe  
                                  690                      695                      700  
 Phe Lys His Ser His Thr Leu Gln Pro Gln Asn Tyr Leu Arg Leu Gly  
 705                                   710                      715                      720  
 Arg Ala Lys Phe Ser Glu Ser Ala Ile Glu Lys Phe Pro Arg Glu Ile  
                                  725                      730                      735  
 Pro Leu Ala Leu Asp Val Gln Val Ser Phe Ser His Ser Asp Asn Arg  
                                  740                      745                      750  
 Met Glu Thr His Tyr Thr Ser Leu Pro Glu Ser Glu Gly Ser Trp Ser  
                                  755                      760                      765

Asn Glu Cys Ile Ala Gly Gly Ile Gly Leu Asp Leu Pro Phe Val Leu  
 770 775 780  
 Ser Asn Pro His Pro Leu Phe Lys Thr Phe Ile Pro Gln Met Lys Val  
 785 790 795 800  
 Glu Met Val Tyr Val Ser Gln Asn Ser Phe Phe Glu Ser Ser Ser Asp  
 805 810 815  
 Gly Arg Gly Phe Ser Ile Gly Arg Leu Leu Asn Leu Ser Ile Pro Val  
 820 825 830  
 Gly Ala Lys Phe Val Gln Gly Asp Ile Gly Asp Ser Tyr Thr Tyr Asp  
 835 840 845  
 Leu Ser Gly Phe Phe Val Ser Asp Val Tyr Arg Asn Asn Pro Gln Ser  
 850 855 860  
 Thr Ala Thr Leu Val Met Ser Pro Asp Ser Trp Lys Ile Arg Gly Gly  
 865 870 875 880  
 Asn Leu Ser Arg Gln Ala Phe Leu Leu Arg Gly Ser Asn Asn Tyr Val  
 885 890 895  
 Tyr Asn Ser Asn Cys Glu Leu Phe Gly His Tyr Ala Met Glu Leu Arg  
 900 905 910  
 Gly Ser Ser Arg Asn Tyr Asn Val Asp Val Gly Thr Lys Leu Arg Phe  
 915 920 925

## (2) INFORMATION FOR SEQ ID NO:3:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2815 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

ATGAAATCGC	AATTTTCCTG	GTTAGTGCTC	TCTTCGACAT	TGGCATGTTT	TACTAGTTGT	60
TCCACTGTTT	TTGCTGCAAC	TGCTGAAAAT	ATAGGCCCTT	CTGATAGCTT	TGACGGAAAGT	120
ACTAACACAG	GCACCTATAC	TCCTAAAAAT	ACGACTACTG	GAATAGACTA	TACTCTGACA	180
GGAGATATAA	CTCTGCAAAA	CCTTGGGGAT	TCGGCAGCTT	TAACGAAGGG	TTGTTTTTCT	240
GACACTACGG	AATCTTTAAG	CTTTGCCGGT	AAGGGGTACT	CACTTTCTTT	TTTAAATATT	300
AAGTCTAGTG	CTGAAGGCGC	AGCACTTTCT	GTTACAACCTG	ATAAAAATCT	GTGCTAACA	360
GGATTTTCGA	GTCTTACTTT	CTTAGCGGCC	CCATCATCGG	TAATCACAAC	CCCCTCAGGA	420
AAAGGTGCAG	TTAAATGTGG	AGGGGATCTT	ACATTTGATA	ACAATGGAAC	TATTTTATTT	480
AAACAAGATT	ACTGTGAGGA	AAATGGCGGA	GCCATTTCTA	CCAAGAATCT	TTCTTTGAAA	540
AACAGCACGG	GATCGATTTT	TTTGAAGGG	AATAAATCGA	GCGCAACAGG	GAAAAAGGT	600
GGGGCTATTT	GTGCTACTGG	TACTGTAGAT	ATTACAAATA	ATACGGCTCC	TACCTCTTC	660
TCGAACAATA	TTGCTGAAGC	TGCAGGTGGA	GCTATAAATA	GCACAGGAAA	CTGTACAATT	720
ACAGGGAATA	CGTCTCTTGT	ATTTTCTGAA	AATAGTGTGA	CAGCGACCGC	AGGAAATGGA	780
GGAGCTCTTT	CTGGAGATGC	CGATGTTACC	ATATCTGGGA	ATCAGAGTGT	AACTTTCTCA	840
GGAAACCAAG	CTGTAGCTAA	TGGCGGAGCC	ATTTATGCTA	AGAAGCTTAC	ACTGGCTTCC	900
GGGGGGGGGG	GGGGTATCTC	CTTTTCTAAC	AATATAGTCC	AAGGTACCAC	TGCAGGTAAT	960
GGTGGAGCCA	TTTCTATACT	GGCAGCTGGA	GAGTGTAGTC	TTTCAGCAGA	AGCAGGGGAC	1020
ATTACCTTCA	ATGGGAATGC	CATTGTTGCA	ACTACACCAC	AAACTACAAA	AAGAAATTCT	1080
ATTGACATAG	GATCTACTGC	AAAGATCACG	AATTTACGTG	CAATATCTGG	GCATAGCATC	1140
TTTTTCTACG	ATCCGATTAC	TGCTAATACG	GCTGCGGATT	CTACAGATAC	TTTAAATCTC	1200
AATAAGGCTG	ATGCAGGTAA	TAGTACAGAT	TATAGTGGGT	CGATTGTTTT	TTCTGGTGAA	1260

AAGCTCTCTG AAGATGAAGC AAAAGTTGCA GACAACCTCA CTTCTACGCT GAAGCAGCCT 1320  
 GTAACCTCTAA CTGCAGGAAA TTTAGTACTT AAACGTGGTG TCACTCTCGA TACGAAAGGC 1380  
 TTTACTCAGA CCGCGGGTTC CTCTGTTATT ATGGATGCGG GCACAACGTT AAAAGCAAGT 1440  
 ACAGAGGAGG TCACTTTAAC AGGTCTTTCC ATTCTGTAG ACTCTTTAGG CGAGGGTAAG 1500  
 AAAGTTGTAA TTGCTGCTTC TGCAGCAAGT AAAAATGTAG CCCTTAGTGG TCCGATTCTT 1560  
 CTTTTGGATA ACCAAGGGAA TGCTTATGAA AATCAGCACT TAGGAAAAAC TCAAGACTTT 1620  
 TCATTTGTGC AGCTCTCTGC TCTGGGTACT GCAACAATA CAGATGTTCC AGCGGTTCCT 1680  
 ACAGTAGCAA CTCCTACGCA CTATGGGTAT CAAGGTACTT GGGGAATGAC TTGGGTGTAT 1740  
 GATACCGCAA GCACTCCAAA GACTAAGACA GCGACATTAG CTTGGACCAA TACAGGCTAC 1800  
 CTTCCGAATC CTGAGCGTCA AGGACCTTTA GTTCCTAATA GCCTTTGGGG ATCTTTTCA 1860  
 GACATCCAAG CGATTCAAGG TGTATAGAG AGAAGTGCTT TGACTCTTG TTCAGATCGA 1920  
 GGCTTCTGGG CTGCGGGAGT CGCCAATTTT TTAGATAAAG ATAAGAAAGG GGAAAAACGC 1980  
 AAATACCGTC ATAAATCTGG TGGATATGCT ATCGGAGGTG CAGCGCAAAC TTGTTCTGAA 2040  
 AACTTAATTA GCTTTGCCCTT TTGCCAACTC TTTGGTAGCG ATAAAGATTT CTAGTCGCT 2100  
 AAAAATCATA CTGATACCTA TGCAGGAGCC TTCTATATCC AACACATTAC AGAATGTAGT 2160  
 GGGTTCATAG GTTGTCTCTT AGATAAACTT CCTGGCTCTT GGAGTCATAA ACCCCTCGTT 2220  
 TTAGAAGGGC AGCTCGCTTA TAGCCACGTC AGTAATGATC TGAAGACAAA GTATACTGCG 2280  
 TATCCTGAGG TGAAAGGFTT TTGGGGGAAT AATGCTTTTA ACATGATGTT GGGAGCTTCT 2340  
 TCTCATTCTT ATCTGAATA CCTGCATTGT TTTGATACCT ATGCTCCATA CATCAAACCTG 2400  
 AATCTGACCT ATATACGTCA GGACAGCTTC TCGGAGAAAG GTACAGAAGG AAGATCTTTT 2460  
 GATGACAGCA ACCTCTTCAA TTTATCTTTG CCTATAGGGG TGAAGTTTGA GAAGTTCTCT 2520  
 GATTGTAATG ACTTTTCTTA TGATCTGACT TTATCCTATG TTCTTGATCT TATCCGCAAT 2580  
 GATCCCAAAT GCACTACAGC ACTTGTATC AGCGGAGCCT CTTGGGAAAC TTATGCCAAT 2640  
 AACTTAGCAC GACAGGCCCT GCAAGTGCGT GCAGGCAGTC ACTACGCCCT CTCTCCTATG 2700  
 TTTGAAGTGC TCGGCCAGTT TGTCTTTGAA GTTCGTGGAT CCTCACGGAT TTATAATGTA 2760  
 GATCTTGGGG GTAAGTTCCA ATTCTAGGAG CGTCTCTCAT GTCTCAGAAA TTCTG 2815

## (2) INFORMATION FOR SEQ ID NO:4:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 928 amino acids  
 (B) TYPE: amino acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

Met Lys Ser Gln Phe Ser Trp Leu Val Leu Ser Ser Thr Leu Ala Cys  
 1 5 10 15  
 Phe Thr Ser Cys Ser Thr Val Phe Ala Thr Ala Glu Asn Ile Gly  
 20 25 30  
 Pro Ser Asp Ser Phe Asp Gly Ser Thr Asn Thr Gly Thr Tyr Thr Pro  
 35 40 45  
 Lys Asn Thr Thr Thr Gly Ile Asp Tyr Thr Leu Thr Gly Asp Ile Thr  
 50 55 60  
 Leu Gln Asn Leu Gly Asp Ser Ala Ala Leu Thr Lys Gly Cys Phe Ser  
 65 70 75 80  
 Asp Thr Thr Glu Ser Leu Ser Phe Ala Gly Lys Gly Tyr Ser Leu Ser  
 85 90 95  
 Phe Leu Asn Ile Lys Ser Ser Ala Glu Gly Ala Ala Leu Ser Val Thr  
 100 105 110  
 Thr Asp Lys Asn Leu Ser Leu Thr Gly Phe Ser Ser Leu Thr Phe Leu  
 115 120 125  
 Ala Ala Pro Ser Ser Val Ile Thr Thr Pro Ser Gly Lys Gly Ala Val  
 130 135 140



Lys Cys Gly Gly Asp Leu Thr Phe Asp Asn Asn Gly Thr Ile Leu Phe  
 145 150 155 160  
 Lys Gln Asp Tyr Cys Glu Glu Asn Gly Gly Ala Ile Ser Thr Lys Asn  
 165 170 175  
 Leu Ser Leu Lys Asn Ser Thr Gly Ser Ile Ser Phe Glu Gly Asn Lys  
 180 185 190  
 Ser Ser Ala Thr Gly Lys Lys Gly Gly Ala Ile Cys Ala Thr Gly Thr  
 195 200 205  
 Val Asp Ile Thr Asn Asn Thr Ala Pro Thr Leu Phe Ser Asn Asn Ile  
 210 215 220  
 Ala Glu Ala Ala Gly Gly Ala Ile Asn Ser Thr Gly Asn Cys Thr Ile  
 225 230 235 240  
 Thr Gly Asn Thr Ser Leu Val Phe Ser Glu Asn Ser Val Thr Ala Thr  
 245 250 255  
 Ala Gly Asn Gly Gly Ala Leu Ser Gly Asp Ala Asp Val Thr Ile Ser  
 260 265 270  
 Gly Asn Gln Ser Val Thr Phe Ser Gly Asn Gln Ala Val Ala Asn Gly  
 275 280 285  
 Gly Ala Ile Tyr Ala Lys Lys Leu Thr Leu Ala Ser Gly Gly Gly Gly  
 290 295 300  
 Gly Ile Ser Phe Ser Asn Asn Ile Val Gln Gly Thr Thr Ala Gly Asn  
 305 310 315 320  
 Gly Gly Ala Ile Ser Ile Leu Ala Ala Gly Glu Cys Ser Leu Ser Ala  
 325 330 335  
 Glu Ala Gly Asp Ile Thr Phe Asn Gly Asn Ala Ile Val Ala Thr Thr  
 340 345 350  
 Pro Gln Thr Thr Lys Arg Asn Ser Ile Asp Ile Gly Ser Thr Ala Lys  
 355 360 365  
 Ile Thr Asn Leu Arg Ala Ile Ser Gly His Ser Ile Phe Phe Tyr Asp  
 370 375 380  
 Pro Ile Thr Ala Asn Thr Ala Ala Asp Ser Thr Asp Thr Leu Asn Leu  
 385 390 395 400  
 Asn Lys Ala Asp Ala Gly Asn Ser Thr Asp Tyr Ser Gly Ser Ile Val  
 405 410 415  
 Phe Ser Gly Glu Lys Leu Ser Glu Asp Glu Ala Lys Val Ala Asp Asn  
 420 425 430  
 Leu Thr Ser Thr Leu Lys Gln Pro Val Thr Leu Thr Ala Gly Asn Leu  
 435 440 445  
 Val Leu Lys Arg Gly Val Thr Leu Asp Thr Lys Gly Phe Thr Gln Thr  
 450 455 460  
 Ala Gly Ser Ser Val Ile Met Asp Ala Gly Thr Thr Leu Lys Ala Ser  
 465 470 475 480  
 Thr Glu Glu Val Thr Leu Thr Gly Leu Ser Ile Pro Val Asp Ser Leu  
 485 490 495  
 Gly Glu Gly Lys Lys Val Val Ile Ala Ala Ser Ala Ala Ser Lys Asn  
 500 505 510  
 Val Ala Leu Ser Gly Pro Ile Leu Leu Leu Asp Asn Gln Gly Asn Ala  
 515 520 525  
 Tyr Glu Asn His Asp Leu Gly Lys Thr Gln Asp Phe Ser Phe Val Gln  
 530 535 540  
 Leu Ser Ala Leu Gly Thr Ala Thr Thr Thr Asp Val Pro Ala Val Pro  
 545 550 555 560  
 Thr Val Ala Thr Pro Thr His Tyr Gly Tyr Gln Gly Thr Trp Gly Met  
 565 570 575  
 Thr Trp Val Asp Asp Thr Ala Ser Thr Pro Lys Thr Lys Thr Ala Thr  
 580 585 590  
 Leu Ala Trp Thr Asn Thr Gly Tyr Leu Pro Asn Pro Glu Arg Gln Gly

595                      600                      605  
 Pro Leu Val Pro Asn Ser Leu Trp Gly Ser Phe Ser Asp Ile Gln Ala  
 610                      615                      620  
 Ile Gln Gly Val Ile Glu Arg Ser Ala Leu Thr Leu Cys Ser Asp Arg  
 625                      630                      635                      640  
 Gly Phe Trp Ala Ala Gly Val Ala Asn Phe Leu Asp Lys Asp Lys Lys  
 645                      650                      655  
 Gly Glu Lys Arg Lys Tyr Arg His Lys Ser Gly Gly Tyr Ala Ile Gly  
 660                      665                      670  
 Gly Ala Ala Gln Thr Cys Ser Glu Asn Leu Ile Ser Phe Ala Phe Cys  
 675                      680                      685  
 Gln Leu Phe Gly Ser Asp Lys Asp Phe Leu Val Ala Lys Asn His Thr  
 690                      695                      700  
 Asp Thr Tyr Ala Gly Ala Phe Tyr Ile Gln His Ile Thr Glu Cys Ser  
 705                      710                      715                      720  
 Gly Phe Ile Gly Cys Leu Leu Asp Lys Leu Pro Gly Ser Trp Ser His  
 725                      730                      735  
 Lys Pro Leu Val Leu Glu Gly Gln Leu Ala Tyr Ser His Val Ser Asn  
 740                      745                      750  
 Asp Leu Lys Thr Lys Tyr Thr Ala Tyr Pro Glu Val Lys Gly Ser Trp  
 755                      760                      765  
 Gly Asn Asn Ala Phe Asn Met Met Leu Gly Ala Ser Ser His Ser Tyr  
 770                      775                      780  
 Pro Glu Tyr Leu His Cys Phe Asp Thr Tyr Ala Pro Tyr Ile Lys Leu  
 785                      790                      795                      800  
 Asn Leu Thr Tyr Ile Arg Gln Asp Ser Phe Ser Glu Lys Gly Thr Glu  
 805                      810                      815  
 Gly Arg Ser Phe Asp Asp Ser Asn Leu Phe Asn Leu Ser Leu Pro Ile  
 820                      825                      830  
 Gly Val Lys Phe Glu Lys Phe Ser Asp Cys Asn Asp Phe Ser Tyr Asp  
 835                      840                      845  
 Leu Thr Leu Ser Tyr Val Pro Asp Leu Ile Arg Asn Asp Pro Lys Cys  
 850                      855                      860  
 Thr Thr Ala Leu Val Ile Ser Gly Ala Ser Trp Glu Thr Tyr Ala Asn  
 865                      870                      875                      880  
 Asn Leu Ala Arg Gln Ala Leu Gln Val Arg Ala Gly Ser His Tyr Ala  
 885                      890                      895  
 Phe Ser Pro Met Phe Glu Val Leu Gly Gln Phe Val Phe Glu Val Arg  
 900                      905                      910  
 Gly Ser Ser Arg Ile Tyr Asn Val Asp Leu Gly Gly Lys Phe Gln Phe  
 915                      920                      925

## (2) INFORMATION FOR SEQ ID NO:5:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 3052 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

ATGCGATTTT	CGCTCTGCGG	ATTTCTCTA	GTTCCTTTCTT	TAACATTGCT	CTCAGTCTTC	60
GACACTTCTT	TGAGTGCTAC	TACGATTTCT	TTAACCCAG	AAGATAGTTT	TCATGGAGAT	120
AGTCAGAATG	CAGAACGTTT	TTATAATGTT	CAAGCTGGGG	ATGTCTATAG	CCTTACTGGT	180

GATGTCTCAA TATCTAACGT CGATAACTCT GCATTAAATA AAGCCTGCTT CAATGTGACC 240  
 TCAGGAAGTG TGACGTTTCG AGGAAATCAT CATGGGTTAT ATTTTAATAA TATTTCCCTCA 300  
 GGAAGTACAA AGGAAGGGGC TGTACTTTGT TGCCAAGATC CTCAAGCAAC GGCACGTTTT 360  
 TCTGGGTTCT CCACGCTCTC TTTTATTCAG AGCCCCGGAG ATATTAAAGA ACAGGGATGT 420  
 CTCTATTCAA AAAATGCACT TATGCTCTTA AACAATTATG TAGTGCGTTT TGAACAAAAC 480  
 CAAAGTAAGA CTAAAGGCGG AGCTATTAGT GGGGCGAATG TTACTATAGT AGGCAACTAC 540  
 GATTCCGTCT CTTTCTATCA GAATGCAGCC ACTTTTGGAG GTGCTATCCA TTCTTCAGGT 600  
 CCCCTACAGA TTGCAGTAAA TCAGGCAGAG ATAAGATTTG CACAAAATAC TGCCAAGAAT 660  
 GGTTCTGGAG GGGCTTTGTA CTCCGATGGT GATATTGATA TTGATCAGAA TGCTTATGTT 720  
 CTATTTGAG AAAATGAGC ATTGACTACT GCTATAGGTA AGGGAGGGGC TGTCTGTTGT 780  
 CTTCCCACTT CAGGAAGTAG TACTCCAGTT CCTATTGTGA CTTTCTCTGA CAATAAACAG 840  
 TTAGTCTTTG AAAGAAACCA TTCCATAATG GGTGGCGGAG CCATTTATGC TAGGAAACTT 900  
 AGCATCTCTT CAGGAGGTCC TACTCTATTT ATCAATAATA TATCATATGC AAATTCGCAA 960  
 AATTTAGGTG GAGCTATTGC CATTGATACT GGAGGGGAGA TCAGTTTATC AGCAGAGAAA 1020  
 GGAACAATTA CATTCCAAGG AAACCGGACG AGCTTACCGT TTTTGAATGG CATCCATCTT 1080  
 TTACAAAATG CTAAATTCCT GAAATTACAG GCGAGAAATG GATGCTCTAT AGAATTTTAT 1140  
 GATCCTATTA CTTCTGAAGC AGATGGGTCT ACCCAATTGA ATATCAACGG AGATCCTAAA 1200  
 AATAAAGAGT ACACAGGGAC CATACTCTTT TCTGGAGAAA AGAGTCTAGC AAACGATCCT 1260  
 AGGGATTTTA AATCTACAAT CCCTCAGAAC GTCAACCTGT CTGCAGGATA CTTAGTTATT 1320  
 AAAGAGGGGG CCGAAGTCAC AGTTTCAAAA TTCACGCAGT CTCCAGGATC GCATTTAGTT 1380  
 TTAGATTTAG GAACCAAACT GATAGCCTCT AAGGAAGACA TTGCCATCAC AGGCCTCGCG 1440  
 ATAGATATAG ATAGCTTAAG CTCATCCTCA ACAGCAGCTG TTATTAAAGC AAACACCGCA 1500  
 AATAACAGA TATCCGTGAC GGACTCTATA GAACTTATCT CGCCTACTGG CAATGCCTAT 1560  
 GAAGATCTCA GAATGAGAAA TTCACAGAG TCCCTCTGC TCTCTTTAGA GCCTGGAGCC 1620  
 GGGGGTAGTG TGACTGTAAC TGCTGGAGAT TTCCTACCGG TAAGTCCCCA TTATGGTTTT 1680  
 CAAGGCAATT GGAAATTAGC TTGGACAGGA ACTGGAAACA AAGTTGGAGA ATTCTTCTGG 1740  
 GATAAAATAA ATTATAAGCC TAGACCTGAA AAAGAAGGAA ATTTAGTTCC TAATATCTTG 1800  
 TGGGGGAATG CTGTAAATGT CAGATCCTTA ATGCAGGTTT AAGAGACCCA TGCATCGAGC 1860  
 TTACAGACAG ATCGAGGGCT GTGGATCGAT GGAATTGGGA ATTTCTTCCA TGTATCTGCC 1920  
 TCCGAAGACA ATATAAGGTA CCGTCATAAC AGCGGTGGAT ATGTTCTATC TGTAATAAAT 1980  
 GAGATCACAC CTAAGCACTA TACTTCGATG GCATTTTCCC AACTCTTTAG TAGAGACAAG 2040  
 GACTATGCGG TTTCCAACAA CGAATACAGA ATGTATTTAG GATCGTATCT CTATCAATAT 2100  
 ACAACCTCCC TAGGGAATAT TTTCCGTTAT GCTTCGCGTA ACCCTAATGT AAACGTCGGG 2160  
 ATTCTCTCAA GAAGGTTTCT TCAAAATCCT CTTATGATTT TTCATTTTTT GTGTGCTTAT 2220  
 GGTCAATGCC CCAATGATAT GAAAACAGAC TACGCAAAT TCCCTATGGT GAAAAACAGC 2280  
 TGGAGAAACA ATTGTTGGGC TATAGAGTGC GGAGGGAGCA TGCCTCTATT GGTATTTGAG 2340  
 AACGGAAGAC TTTTCCAAGG TGCCATCCCA TTTATGAAAC TACAATTAGT TTATGCTTAT 2400  
 CAGGGAGATT TCAAAGAGAC GACTGCAGAT GGCCGTAGAT TTAGTAATGG GAGTTTAACA 2460  
 TCGATTTCTG TACCTCTAGG CATAAGCTTT GAGAAGCTGG CACTTTCTCA GGATGTACTC 2520  
 TATGACTTTA GTTCTCTCTA TATTCCTGAT ATTTTCCGTA AGGATCCCTC ATGTGAAGCT 2580  
 GCTCTGGTGA TTAGCGGAGA CTCCTGGCTT GTTCCGGCAG CACACGTATC AAGACATGCT 2640  
 TTTGTAGGGA GTGGAACGGG TCGGTATCAC TTTAACGACT ATACTGAGCT CTTATGTGCA 2700  
 GGAAGTATAG AATGCCGCC CCATGCTAGG AATTATAATA TAACTGTGG AAGCAAATTT 2760  
 CGTTTTTAGA AGGTTTCCAT TGCCTGTGTG GTTCCGGATC TTAACATAA ATCCTGGACT 2820  
 ATGGATCATA GGCATTGGGT TTCTCGAACT TGTGTGGAGA ATAAAGACAT TTTATATGCA 2880  
 TAACGGAATA CTCGTATCAC CTCAGCCCT AGAGACATTC TTTAGGGGTT CTTTATTTGT 2940  
 CTAACTTCG TATTTTATCG AGAATCCTTT ACGTTCTTGG TTTGCTTGTG TCCGAGGAGT 3000  
 TCTCTAACGA ATCATAGGGA TTCCAGGGTT CTGTTCTTGG AGTCCTTTGG CA 3052

## (2) INFORMATION FOR SEQ ID NO:6:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 922 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

```

Met Arg Phe Ser Leu Cys Gly Phe Pro Leu Val Phe Ser Leu Thr Leu
 1           5           10           15
Leu Ser Val Phe Asp Thr Ser Leu Ser Ala Thr Thr Ile Ser Leu Thr
           20           25           30
Pro Glu Asp Ser Phe His Gly Asp Ser Gln Asn Ala Glu Arg Ser Tyr
           35           40           45
Asn Val Gln Ala Gly Asp Val Tyr Ser Leu Thr Gly Asp Val Ser Ile
           50           55           60
Ser Asn Val Asp Asn Ser Ala Leu Asn Lys Ala Cys Phe Asn Val Thr
65           70           75           80
Ser Gly Ser Val Thr Phe Ala Gly Asn His His Gly Leu Tyr Phe Asn
           85           90           95
Asn Ile Ser Ser Gly Thr Thr Lys Glu Gly Ala Val Leu Cys Cys Gln
           100           105           110
Asp Pro Gln Ala Thr Ala Arg Phe Ser Gly Phe Ser Thr Leu Ser Phe
           115           120           125
Ile Gln Ser Pro Gly Asp Ile Lys Glu Gln Gly Cys Leu Tyr Ser Lys
           130           135           140
Asn Ala Leu Met Leu Leu Asn Asn Tyr Val Val Arg Phe Glu Gln Asn
145           150           155           160
Gln Ser Lys Thr Lys Gly Gly Ala Ile Ser Gly Ala Asn Val Thr Ile
           165           170           175
Val Gly Asn Tyr Asp Ser Val Ser Phe Tyr Gln Asn Ala Ala Thr Phe
           180           185           190
Gly Gly Ala Ile His Ser Ser Gly Pro Leu Gln Ile Ala Val Asn Gln
           195           200           205
Ala Glu Ile Arg Phe Ala Gln Asn Thr Ala Lys Asn Gly Ser Gly Gly
210           215           220
Ala Leu Tyr Ser Asp Gly Asp Ile Asp Ile Asp Gln Asn Ala Tyr Val
225           230           235           240
Leu Phe Arg Glu Asn Glu Ala Leu Thr Thr Ala Ile Gly Lys Gly Gly
           245           250           255
Ala Val Cys Cys Leu Pro Thr Ser Gly Ser Ser Thr Pro Val Pro Ile
           260           265           270
Val Thr Phe Ser Asp Asn Lys Gln Leu Val Phe Glu Arg Asn His Ser
           275           280           285
Ile Met Gly Gly Gly Ala Ile Tyr Ala Arg Lys Leu Ser Ile Ser Ser
           290           295           300
Gly Gly Pro Thr Leu Phe Ile Asn Asn Ile Ser Tyr Ala Asn Ser Gln
305           310           315           320
Asn Leu Gly Gly Ala Ile Ala Ile Asp Thr Gly Gly Glu Ile Ser Leu
           325           330           335
Ser Ala Glu Lys Gly Thr Ile Thr Phe Gln Gly Asn Arg Thr Ser Leu
           340           345           350
Pro Phe Leu Asn Gly Ile His Leu Leu Gln Asn Ala Lys Phe Leu Lys
           355           360           365
Leu Gln Ala Arg Asn Gly Cys Ser Ile Glu Phe Tyr Asp Pro Ile Thr
           370           375           380
Ser Glu Ala Asp Gly Ser Thr Gln Leu Asn Ile Asn Gly Asp Pro Lys
385           390           395           400
Asn Lys Glu Tyr Thr Gly Thr Ile Leu Phe Ser Gly Glu Lys Ser Leu
           405           410           415
Ala Asn Asp Pro Arg Asp Phe Lys Ser Thr Ile Pro Gln Asn Val Asn

```

420 425 430  
 Leu Ser Ala Gly Tyr Leu Val Ile Lys Glu Gly Ala Glu Val Thr Val  
 435 440 445  
 Ser Lys Phe Thr Gln Ser Pro Gly Ser His Leu Val Leu Asp Leu Gly  
 450 455 460  
 Thr Lys Leu Ile Ala Ser Lys Glu Asp Ile Ala Ile Thr Gly Leu Ala  
 465 470 475 480  
 Ile Asp Ile Asp Ser Leu Ser Ser Ser Ser Thr Ala Ala Val Ile Lys  
 485 490 495  
 Ala Asn Thr Ala Asn Lys Gln Ile Ser Val Thr Asp Ser Ile Glu Leu  
 500 505 510  
 Ile Ser Pro Thr Gly Asn Ala Tyr Glu Asp Leu Arg Met Arg Asn Ser  
 515 520 525  
 Gln Thr Phe Pro Leu Leu Ser Leu Glu Pro Gly Ala Gly Gly Ser Val  
 530 535 540  
 Thr Val Thr Ala Gly Asp Phe Leu Pro Val Ser Pro His Tyr Gly Phe  
 545 550 555 560  
 Gln Gly Asn Trp Lys Leu Ala Trp Thr Gly Thr Gly Asn Lys Val Gly  
 565 570 575  
 Glu Phe Phe Trp Asp Lys Ile Asn Tyr Lys Pro Arg Pro Glu Lys Glu  
 580 585 590  
 Gly Asn Leu Val Pro Asn Ile Leu Trp Gly Asn Ala Val Asn Val Arg  
 595 600 605  
 Ser Leu Met Gln Val Gln Glu Thr His Ala Ser Ser Leu Gln Thr Asp  
 610 615 620  
 Arg Gly Leu Trp Ile Asp Gly Ile Gly Asn Phe Phe His Val Ser Ala  
 625 630 635 640  
 Ser Glu Asp Asn Ile Arg Tyr Arg His Asn Ser Gly Gly Tyr Val Leu  
 645 650 655  
 Ser Val Asn Asn Glu Ile Thr Pro Lys His Tyr Thr Ser Met Ala Phe  
 660 665 670  
 Ser Gln Leu Phe Ser Arg Asp Lys Asp Tyr Ala Val Ser Asn Asn Glu  
 675 680 685  
 Tyr Arg Met Tyr Leu Gly Ser Tyr Leu Tyr Gln Tyr Thr Thr Ser Leu  
 690 695 700  
 Gly Asn Ile Phe Arg Tyr Ala Ser Arg Asn Pro Asn Val Asn Val Gly  
 705 710 715 720  
 Ile Leu Ser Arg Arg Phe Leu Gln Asn Pro Leu Met Ile Phe His Phe  
 725 730 735  
 Leu Cys Ala Tyr Gly His Ala Thr Asn Asp Met Lys Thr Asp Tyr Ala  
 740 745 750  
 Asn Phe Pro Met Val Lys Asn Ser Trp Arg Asn Asn Cys Trp Ala Ile  
 755 760 765  
 Glu Cys Gly Gly Ser Met Pro Leu Leu Val Phe Glu Asn Gly Arg Leu  
 770 775 780  
 Phe Gln Gly Ala Ile Pro Phe Met Lys Leu Gln Leu Val Tyr Ala Tyr  
 785 790 795 800  
 Gln Gly Asp Phe Lys Glu Thr Thr Ala Asp Gly Arg Arg Phe Ser Asn  
 805 810 815  
 Gly Ser Leu Thr Ser Ile Ser Val Pro Leu Gly Ile Arg Phe Glu Lys  
 820 825 830  
 Leu Ala Leu Ser Gln Asp Val Leu Tyr Asp Phe Ser Phe Ser Tyr Ile  
 835 840 845  
 Pro Asp Ile Phe Arg Lys Asp Pro Ser Cys Glu Ala Ala Leu Val Ile  
 850 855 860  
 Ser Gly Asp Ser Trp Leu Val Pro Ala Ala His Val Ser Arg His Ala  
 865 870 875 880

(A) LENGTH: 2526 base pairs  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEO ID NO:7:

ATGAAGATTTC	CACCTCCGCTT	TTTATTGATA	TCATTAGTAC	CTACGCTTTC	TATGTCGAAT	60
TTATTAGGAG	CTGCTACTAC	CGAAGAGCTA	TCGGCTAGCA	ATAGCTTCGA	TGGAAC TACA	120
TCAACAACAA	GCTTTTCTAG	TAAAACATCA	TCGGCTACAG	ATGGCACC AA	TTATGTTTTT	180
AAAGATTCTG	TAGTTATAGA	AAATGTACCC	AAAACAGGGG	AAACTCAGTC	TACTAGTTGT	240
TTTAAAAATG	ACGCTGCAGC	TGGAGATCTA	AATTTCTTAG	GAGGGGGATT	TTCTTTTACA	300
TTTAGCAATA	TCGATGCAAC	CACGGCTTCT	GGAGCTGCTA	TTGGAAGTGA	AGCAGCTAAT	360
AAGACAGTCA	CGTTATCAGG	ATTTTCGGCA	CTTTCTTTTC	TTAAATCCCC	AGCAAGTACA	420
GTGACTAATG	GATTGGGAGC	TATCAATGTT	AAAGGGAATT	TAAGCCTATT	GGATAATGAT	480
AAGGTATTGA	TTCAGGACAA	TTTCTCAACA	GGAGATGGCG	GAGCAATTAA	TTGTGCAGGC	540
TCCTTGAAGA	TCGCAAAACA	TAAGTCCCTT	TCCTTTTATTG	GAAATAGTTT	TTCAACACGT	600
GGCGGAGCGA	TTTCATACCA	AAACCTCACA	CTATCTTCTG	TGTGGGAAAC	TCTATTTTCAG	660
GGGAATACAG	CGCCTACGGC	TGCTGGTAAA	GGAGGTGCTA	TCGCGATTGC	AGACTCTGGC	720
ACCCTATCCA	TTTCTGGAGA	CAGTGGCGAC	ATTATCTTTG	AAGGCAATAC	GATAGGAGCT	780
ACAGGAACCG	TCTCTCATAG	TGCTATTGAT	TTAGGAACTA	GCGCTAAGAT	AACTGCGTTA	840
CGTGCTGCGC	AAGGACATAC	GATATACTTT	TATGATCCGA	TTACTGTAAC	AGGATCGACA	900
TCTGTTGCTG	ATGCTCTCAA	TATTAATAGC	CCTGATACTG	GAGATAACAA	AGAGTATACG	960
GGAACCATAG	TCTTTTCTGG	AGAGAAGCTC	ACGGAGGCAG	AAGCTAAAGA	TGAGAAGAAC	1020
CGCACTTCTA	AATTACTTCA	AAATGTTGCT	TTTAAAAATG	GGACTGTAGT	TTTAAAAGGT	1080
GATGTCGTTT	TAAGTGCGAA	CGGTTTCTCT	CAGGATGCAA	ACTCTAAGTT	GATTATGGAT	1140
TTAGGGACGT	CGTTGGTTGC	AAACACCGAA	AGTATCGAGT	TAACGAATTT	GGAAATTAAT	1200
ATAGACTCTC	TCAGGAACGG	GAAAAAGATA	AAACTCAGTG	CTGCCACAGC	TCAGAAAGAT	1260
ATTCGTATAG	ATCGTCCGTG	TGTACTGGCA	ATTAGCGTAG	AGAGTTTTTA	TCAAAATGGC	1320
TTTTTGAATG	AGGACCATTC	CTATAGTGGG	ATTCTTGATG	TAGATGCTGG	GAAAGACATC	1380
GTGATTTCTG	CAGATTCTCG	CAGTATAAAT	GCTGTACAAT	CTCCGTATGG	CTATCAGGGA	1440
AAGTGGACAA	TCAATTGGTC	TACTGATGAT	AAGAAAGCTA	CGGTTTCTTG	GGCAAAGCAA	1500
AGTTTTTAATC	CCACTGCTGA	GCAGGAGGCT	CCGTTAGTTC	CTAATCTTCT	TTGGGGTTCT	1560
TTTATAGATG	TTCGTCCCTT	CCAAAATTTT	ATAGAGCTAG	GTACTGAAGG	TGCTCCCTTAC	1620
GAAAAGAGAT	TTTGGGTTGC	AGGCATTTCC	AATGTTTTGC	ATAGGAGCGG	TCGTGAAAT	1680
CAAAGGAAAT	TCCGT CATGT	GAGTGGAGGT	GCTGTAGTAG	GTGCTAGCAC	GAGGATGCCG	1740
GGTGGTGATA	CCTGTCTCT	GGGTTTTGCT	CAGCTCTTTG	CGCGTGACAA	AGACTACTTT	1800
ATGAATACCA	ATTCGCAAA	GACCTACGCA	GGATCTTTAC	GTTTGCAGCA	CGATGCTTCC	1860
CTATACTCTG	TGGTGAGTAT	CCTTTTAGGA	GAGGGAGGAC	TCCGCGAGAT	CCTGTTGCCT	1920
TATGTTTTCCA	AGACTCTGCC	GTGCTCTTTC	TATGGGCAGC	TTAGCTACGG	CCATACGGAT	1980
CATCGCATGA	AGACCGAGTC	TCTACCCCCC	CCCCCCCCGA	CGCTCTCGAC	GGATCATACT	2040
TCTTTGGGAG	GATATGTCTG	GGCTGGAGAG	CTGGGAACTC	GAGTTGCTGT	TGAAATATACC	2100
AGCGGCAGAG	GATTTTCCG	AGAGTACACT	CCATTTGTAA	AAGTCCAAGC	TGTTTACTCG	2160
GCCCAAGATA	GCTTGTGTGA	ACTAGGAGCT	ATCAGTCTGT	ATTTTAGTGA	TTGCGATCTT	2220
TATAACCTTG	CGATTCCCTCT	TGGAATCAAG	TTAGAGAAAC	GGTTTGCAGA	GCAATATTAT	2280

CATGTTGTAG	CGATGTATTC	TCCAGATGTT	TGTCGTAGTA	ACCCCAAATG	TACGACTACC	2340
CTACTTTCCA	ACCAAGGGAG	TTGGAAGACC	AAAGGTTTCA	ACTTAGCAAG	ACAGGCTGGT	2400
ATTGTTTCAGG	CCTCAGGTTT	TCGATCTTTG	GGAGCTGCAG	CAGAGCTTTT	CGGGAACTTT	2460
GGCTTTGAAT	GGCGGGGATC	TTCTCGTAGC	TATAATGTAG	ATGCGGGTAG	CAAAATCAAA	2520
TTTTAG						2526

## (2) INFORMATION FOR SEQ ID NO:8:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 841 amino acids  
 (B) TYPE: amino acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

Met	Lys	Ile	Pro	Leu	Arg	Phe	Leu	Leu	Ile	Ser	Leu	Val	Pro	Thr	Leu	1	5	10	15
Ser	Met	Ser	Asn	Leu	Leu	Gly	Ala	Ala	Thr	Thr	Glu	Glu	Leu	Ser	Ala	20	25	30	
Ser	Asn	Ser	Phe	Asp	Gly	Thr	Thr	Ser	Thr	Thr	Ser	Phe	Ser	Ser	Lys	35	40	45	
Thr	Ser	Ser	Ala	Thr	Asp	Gly	Thr	Asn	Tyr	Val	Phe	Lys	Asp	Ser	Val	50	55	60	
Val	Ile	Glu	Asn	Val	Pro	Lys	Thr	Gly	Glu	Thr	Gln	Ser	Thr	Ser	Cys	65	70	75	80
Phe	Lys	Asn	Asp	Ala	Ala	Gly	Asp	Leu	Asn	Phe	Leu	Gly	Gly	Gly		85	90	95	
Phe	Ser	Phe	Thr	Phe	Ser	Asn	Ile	Asp	Ala	Thr	Thr	Ala	Ser	Gly	Ala	100	105	110	
Ala	Ile	Gly	Ser	Glu	Ala	Ala	Asn	Lys	Thr	Val	Thr	Leu	Ser	Gly	Phe	115	120	125	
Ser	Ala	Leu	Ser	Phe	Leu	Lys	Ser	Pro	Ala	Ser	Thr	Val	Thr	Asn	Gly	130	135	140	
Leu	Gly	Ala	Ile	Asn	Val	Lys	Gly	Asn	Leu	Ser	Leu	Leu	Asp	Asn	Asp	145	150	155	160
Lys	Val	Leu	Ile	Gln	Asp	Asn	Phe	Ser	Thr	Gly	Asp	Gly	Gly	Ala	Ile	165	170	175	
Asn	Cys	Ala	Gly	Ser	Leu	Lys	Ile	Ala	Asn	Asn	Lys	Ser	Leu	Ser	Phe	180	185	190	
Ile	Gly	Asn	Ser	Ser	Ser	Thr	Arg	Gly	Gly	Ala	Ile	His	Thr	Lys	Asn	195	200	205	
Leu	Thr	Leu	Ser	Ser	Gly	Gly	Glu	Thr	Leu	Phe	Gln	Gly	Asn	Thr	Ala	210	215	220	
Pro	Thr	Ala	Ala	Gly	Lys	Gly	Gly	Ala	Ile	Ala	Ile	Ala	Asp	Ser	Gly	225	230	235	240
Thr	Leu	Ser	Ile	Ser	Gly	Asp	Ser	Gly	Asp	Ile	Ile	Phe	Glu	Gly	Asn	245	250	255	
Thr	Ile	Gly	Ala	Thr	Gly	Thr	Val	Ser	His	Ser	Ala	Ile	Asp	Leu	Gly	260	265	270	
Thr	Ser	Ala	Lys	Ile	Thr	Ala	Leu	Arg	Ala	Ala	Gln	Gly	His	Thr	Ile	275	280	285	
Tyr	Phe	Tyr	Asp	Pro	Ile	Thr	Val	Thr	Gly	Ser	Thr	Ser	Val	Ala	Asp	290	295	300	
Ala	Leu	Asn	Ile	Asn	Ser	Pro	Asp	Thr	Gly	Asp	Asn	Lys	Glu	Tyr	Thr				

305                    310                    315                    320  
 Gly Thr Ile Val Phe Ser Gly Glu Lys Leu Thr Glu Ala Glu Ala Lys  
                                  325                    330                    335  
 Asp Glu Lys Asn Arg Thr Ser Lys Leu Leu Gln Asn Val Ala Phe Lys  
                                  340                    345                    350  
 Asn Gly Thr Val Val Leu Lys Gly Asp Val Val Leu Ser Ala Asn Gly  
                                  355                    360                    365  
 Phe Ser Gln Asp Ala Asn Ser Lys Leu Ile Met Asp Leu Gly Thr Ser  
                                  370                    375                    380  
 Leu Val Ala Asn Thr Glu Ser Ile Glu Leu Thr Asn Leu Glu Ile Asn  
 385                                   390                    395                    400  
 Ile Asp Ser Leu Arg Asn Gly Lys Lys Ile Lys Leu Ser Ala Ala Thr  
                                  405                    410                    415  
 Ala Gln Lys Asp Ile Arg Ile Asp Arg Pro Val Val Leu Ala Ile Ser  
                                  420                    425                    430  
 Asp Glu Ser Phe Tyr Gln Asn Gly Phe Leu Asn Glu Asp His Ser Tyr  
                                  435                    440                    445  
 Asp Gly Ile Leu Glu Leu Asp Ala Gly Lys Asp Ile Val Ile Ser Ala  
                                  450                    455                    460  
 Asp Ser Arg Ser Ile Asn Ala Val Gln Ser Pro Tyr Gly Tyr Gln Gly  
 465                                   470                    475                    480  
 Lys Trp Thr Ile Asn Trp Ser Thr Asp Asp Lys Lys Ala Thr Val Ser  
                                  485                    490                    495  
 Trp Ala Lys Gln Ser Phe Asn Pro Thr Ala Glu Gln Glu Ala Pro Leu  
                                  500                    505                    510  
 Val Pro Asn Leu Leu Trp Gly Ser Phe Ile Asp Val Arg Pro Phe Gln  
                                  515                    520                    525  
 Asn Phe Ile Glu Leu Gly Thr Glu Gly Ala Pro Tyr Glu Lys Arg Phe  
                                  530                    535                    540  
 Trp Val Ala Gly Ile Ser Asn Val Leu His Arg Ser Gly Arg Glu Asn  
 545                                   550                    555                    560  
 Gln Arg Lys Phe Arg His Val Ser Gly Gly Ala Val Val Gly Ala Ser  
                                  565                    570                    575  
 Thr Arg Met Pro Gly Gly Asp Thr Leu Ser Leu Gly Phe Ala Gln Leu  
                                  580                    585                    590  
 Phe Ala Arg Asp Lys Asp Tyr Phe Met Asn Thr Asn Phe Ala Lys Thr  
                                  595                    600                    605  
 Tyr Ala Gly Ser Leu Arg Leu Gln His Asp Ala Ser Leu Tyr Ser Val  
                                  610                    615                    620  
 Val Ser Ile Leu Leu Gly Glu Gly Gly Leu Arg Glu Ile Leu Leu Pro  
 625                                   630                    635                    640  
 Tyr Val Ser Lys Thr Leu Pro Cys Ser Phe Tyr Gly Gln Leu Ser Tyr  
                                  645                    650                    655  
 Gly His Thr Asp His Arg Met Lys Thr Glu Ser Leu Pro Pro Pro Pro  
                                  660                    665                    670  
 Pro Thr Leu Ser Thr Asp His Thr Ser Trp Gly Gly Tyr Val Trp Ala  
                                  675                    680                    685  
 Gly Glu Leu Gly Thr Arg Val Ala Val Glu Asn Thr Ser Gly Arg Gly  
                                  690                    695                    700  
 Phe Phe Arg Glu Tyr Thr Pro Phe Val Lys Val Gln Ala Val Tyr Ser  
 705                                   710                    715                    720  
 Arg Gln Asp Ser Phe Val Glu Leu Gly Ala Ile Ser Arg Asp Phe Ser  
                                  725                    730                    735  
 Asp Ser His Leu Tyr Asn Leu Ala Ile Pro Leu Gly Ile Lys Leu Glu  
                                  740                    745                    750  
 Lys Arg Phe Ala Glu Gln Tyr Tyr His Val Val Ala Met Tyr Ser Pro  
                                  755                    760                    765



Asp Val Cys Arg Ser Asn Pro Lys Cys Thr Thr Thr Leu Leu Ser Asn  
 770 775 780  
 Gln Gly Ser Trp Lys Thr Lys Gly Ser Asn Leu Ala Arg Gln Ala Gly  
 785 790 795 800  
 Ile Val Gln Ala Ser Gly Phe Arg Ser Leu Gly Ala Ala Ala Glu Leu  
 805 810 815  
 Phe Gly Asn Phe Gly Phe Glu Trp Arg Gly Ser Ser Arg Ser Tyr Asn  
 820 825 830  
 Val Asp Ala Gly Ser Lys Ile Lys Phe  
 835 840

## (2) INFORMATION FOR SEQ ID NO:9:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2787 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:

ATGAAGTCTT	CTTCCCAA	GTTTGTATTT	TCTACATTTG	CTATTTTCCC	TTTGTCTATG	60
ATTGCTACCG	AGACAGTTTT	GGATTCAAGT	GCGAGTTTCG	ATGGGAATAA	AAATGGTAAT	120
TTTTCAGTTC	GTGAGAGTCA	GGAAGATGCT	GGAACACCT	ACCTATTTAA	GGGAAATGTC	180
ACTCTAGAAA	ATATTCCTGG	AACAGGCACA	GCAATCACAA	AAAGCTGTTT	TAACAACACT	240
AAGGGCGATT	TGACTTTCAC	AGGTAACGGG	AACTCTCTAT	TGTTCCAAAC	GGTGGATGCA	300
GGGACTGTAG	CAGGGGCTGC	TGTTAACAGC	AGCGTGCTAG	ATAAATCTAC	CACGTTTATA	360
GGGTTTTCTT	CGCTATCTTT	TATTGCGTCT	CCTGGAAGTT	CGATAACTAC	CGGCAAAGGA	420
GCCGTTAGCT	GCTCTACGGG	TAGCTTGAAG	TTTGACAAAA	ATGTCAGTTT	GCTCTTCAGC	480
AAAAACTTTT	CAACGGATAA	TGGCGGTGCT	ATCACCGCAA	AAACTCTTTC	ATTAACAGGG	540
ACTACAATGT	CAGCTCTGTT	TTCTGAAAT	ACCTCCTCAA	AGAAAGGCGG	AGCCATTCAG	600
ACTTCCGATG	CCCTTACCAT	TACTGGAAAC	CAAGGGGAAG	TCTCTTTTTC	TGACAATACT	660
TCTTCGATT	CTGGAGCTGC	AATTTTTACA	GAAGCCTCGG	TGACTATTTT	TAATAATGCT	720
AAAGTTTCCT	TTATTGACAA	TAAGGTCACA	GGAGCGAGCT	CCTCAACAAC	GGGGGATATG	780
TCAGGAGGTG	CTATCTGTGC	TTATAAAACT	AGTACAGATA	CTAAGGTCAC	CCTCACTGGA	840
AATCAGATGT	TACTCTTCAG	CAACAATACA	TCGACAACAG	CGGGAGGAGC	TATCTATGTG	900
AAAAAGCTCG	AACTGGCTTC	CGGAGGACTT	ACCCTATTCA	GTAGAAATAG	TGTCAATGGA	960
GGTACAGCTC	CTAAAGGTGG	AGCCATAGCT	ATCGAAGATA	GTGGGGAATT	GAGTTTATCC	1020
GCCGATAGTG	GTGACATTGT	CTTTTTAGGG	AATACAGTCA	CTTCTACTAC	TCCTGGGACG	1080
AATAGAAGTA	GTATCGACTT	AGGAACGAGT	GCAAAGATGA	CAGCTTTGCG	TTCTGCTGCT	1140
GGTAGAGCCA	TCTACTTCTA	TGATCCCATA	ACTACAGGAT	CTTCCACAAC	AGTTACAGAT	1200
GTCTTAAAAG	TTAATGAGAC	TCCGGCAGAT	TCTGCCTAC	AATATACAGG	GAACATCATC	1260
TTCACAGGAG	AAAAGTTATC	AGAGACAGAG	GCCGCAGATT	CTAAAAATCT	TACTTCGAAG	1320
CTACTACAGC	CTGTAACTCT	TTCAGGAGGT	ACTCTATCTT	TAAAACATGG	AGTGACTCTG	1380
CAGACTCAGG	CATTCACTCA	ACAGGCAGAT	TCTCGTCTCG	AAATGGACGT	AGGAACTACT	1440
CTAGAACCTG	CTGATACTAG	CACCATAAAC	AATTTGGTCA	TTAACATCAG	TTCTATAGAC	1500
CTTGCAAAGA	AGGCAAAAT	AGAAACCAAA	GCTACGTCAA	AAAATCTGAC	TTTATCTGGA	1560
ACCATCACTT	TATTGGACCC	GACGGGCACG	TTTTATGAAA	ATCATAGTTT	AAGAAATCCT	1620
CAGTCCTACG	ACATCTTAGA	GCTCAAAGCT	TCTGGAAGTG	TAACAAGCAC	CGCAGTGACT	1680
CCAGATCCTA	TAATGGGTGA	GAAATTCCAT	TACGGCTATC	AGGGAAGTTG	GGGCCCAATT	1740
GTTTGGGGGA	CAGGGGCTTC	TACGACTGCA	ACCTTCAACT	GGACTAAAAC	TGGCTATATT	1800
CCTAATCCCG	AGCGTATCGG	CTCTTTAGTC	CCTTAAGCT	TATGGAATGC	ATTTATAGAT	1860
ATTAGCTCTC	TCCATTATCT	TATGGAGACT	GCAAACGAAG	GGTTGCAGGG	AGACCGTGCT	1920
TTTTGGTGTG	CTGGATTATC	TAACCTCTTC	CATAAGGATA	GTACAAAAAC	ACGACGCGGG	1980
TTTCGCCATT	TGAGTGGCGG	TTATGTCATA	GGAGGAAACC	TACATACTTG	TTCAGATAAG	2040

ATTCTTAGTG	CTGCATTTTG	TCAGCTCTTT	GGAAGAGATA	GAGACTACTT	TGTAGCTAAG	2100
AATCAAGGTA	CAGTCTACGG	AGGAAGTCTC	TATTACCAGC	ACAACGAAAC	CTATATCTCT	2160
CTTCCTTGCA	AACTACGGCC	TTGTTGCTTG	TCTTATGTTT	CTACAGAGAT	TCCTGTTCTC	2220
TTTTTCAGGAA	ACCTTAGCTA	CACCCATACG	GATAACGATC	TGAAAACCAA	GTATACAACA	2280
TATCCTACTG	TTAAAGGAAG	CTGGGGGAAT	GATAGTTTCG	CTTTAGAATT	CGGTGGAAGA	2340
GCTCCGATTT	GCTTAGATGA	AAGTGCTCTA	TTTGAGCAGT	ACATGCCCTT	CATGAAATTG	2400
CAGTTTGTCT	ATGCACATCA	GGAAGGTTTT	AAAGAACAGG	GAACAGAAGC	TCGTGAATTT	2460
GGAAGTAGCC	GTCCTGTGAA	TCTTGCCTTA	CCTATCGGGA	TCCGATTTGA	TAAGGAATCA	2520
GACTGCCAAG	ATGCAACGTA	CAATCTAACT	CTTGGTTATA	CTGTGGATCT	TGTTTCGTAGT	2580
AACCCCGACT	GTACGACAAC	ACTGCGAATT	AGCGGTGATT	CTTGGAAAAC	CTTCGGTACG	2640
AATTTGGCAA	GACAAGCITT	AGTCCTTCGT	GCAGGGAACC	ATTTTTCGCT	TAACTCAAT	2700
TTTGAAGCCT	TTAGCCAATT	TTCTTTTGAA	TTGCGTGGGT	CATCTCGCAA	TTACAATGTA	2760
GACTTAGGAG	CAAAATACCA	ATTCTAA				2787

## (2) INFORMATION FOR SEQ ID NO:10:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 928 amino acids  
 (B) TYPE: amino acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:

Met	Lys	Ser	Ser	Phe	Pro	Lys	Phe	Val	Phe	Ser	Thr	Phe	Ala	Ile	Phe
1				5				10						15	
Pro	Leu	Ser	Met	Ile	Ala	Thr	Glu	Thr	Val	Leu	Asp	Ser	Ser	Ala	Ser
			20					25					30		
Phe	Asp	Gly	Asn	Lys	Asn	Gly	Asn	Phe	Ser	Val	Arg	Glu	Ser	Gln	Glu
		35					40					45			
Asp	Ala	Gly	Thr	Thr	Tyr	Leu	Phe	Lys	Gly	Asn	Val	Thr	Leu	Glu	Asn
	50					55					60				
Ile	Pro	Gly	Thr	Gly	Thr	Ala	Ile	Thr	Lys	Ser	Cys	Phe	Asn	Asn	Thr
65					70					75				80	
Lys	Gly	Asp	Leu	Thr	Phe	Thr	Gly	Asn	Gly	Asn	Ser	Leu	Leu	Phe	Gln
			85					90					95		
Thr	Val	Asp	Ala	Gly	Thr	Val	Ala	Gly	Ala	Ala	Val	Asn	Ser	Ser	Val
		100						105					110		
Val	Asp	Lys	Ser	Thr	Thr	Phe	Ile	Gly	Phe	Ser	Ser	Leu	Ser	Phe	Ile
	115						120					125			
Ala	Ser	Pro	Gly	Ser	Ser	Ile	Thr	Thr	Gly	Lys	Gly	Ala	Val	Ser	Cys
	130					135					140				
Ser	Thr	Gly	Ser	Leu	Lys	Phe	Asp	Lys	Asn	Val	Ser	Leu	Leu	Phe	Ser
145					150					155				160	
Lys	Asn	Phe	Ser	Thr	Asp	Asn	Gly	Gly	Ala	Ile	Thr	Ala	Lys	Thr	Leu
			165					170					175		
Ser	Leu	Thr	Gly	Thr	Thr	Met	Ser	Ala	Leu	Phe	Ser	Glu	Asn	Thr	Ser
		180						185					190		
Ser	Lys	Lys	Gly	Gly	Ala	Ile	Gln	Thr	Ser	Asp	Ala	Leu	Thr	Ile	Thr
	195					200						205			
Gly	Asn	Gln	Gly	Glu	Val	Ser	Phe	Ser	Asp	Asn	Thr	Ser	Ser	Asp	Ser
	210					215					220				
Gly	Ala	Ala	Ile	Phe	Thr	Glu	Ala	Ser	Val	Thr	Ile	Ser	Asn	Asn	Ala
225					230					235				240	
Lys	Val	Ser	Phe	Ile	Asp	Asn	Lys	Val	Thr	Gly	Ala	Ser	Ser	Ser	Thr

245 250 255  
 Thr Gly Asp Met Ser Gly Gly Ala Ile Cys Ala Tyr Lys Thr Ser Thr  
 260 265 270  
 Asp Thr Lys Val Thr Leu Thr Gly Asn Gln Met Leu Leu Phe Ser Asn  
 275 280 285  
 Asn Thr Ser Thr Thr Ala Gly Gly Ala Ile Tyr Val Lys Lys Leu Glu  
 290 295 300  
 Leu Ala Ser Gly Gly Leu Thr Leu Phe Ser Arg Asn Ser Val Asn Gly  
 305 310 315 320  
 Gly Thr Ala Pro Lys Gly Gly Ala Ile Ala Ile Glu Asp Ser Gly Glu  
 325 330 335  
 Leu Ser Leu Ser Ala Asp Ser Gly Asp Ile Val Phe Leu Gly Asn Thr  
 340 345 350  
 Val Thr Ser Thr Thr Pro Gly Thr Asn Arg Ser Ser Ile Asp Leu Gly  
 355 360 365  
 Thr Ser Ala Lys Met Thr Ala Leu Arg Ser Ala Ala Gly Arg Ala Ile  
 370 375 380  
 Tyr Phe Tyr Asp Pro Ile Thr Thr Gly Ser Ser Thr Thr Val Thr Asp  
 385 390 395 400  
 Val Leu Lys Val Asn Glu Thr Pro Ala Asp Ser Ala Leu Gln Tyr Thr  
 405 410 415  
 Gly Asn Ile Ile Phe Thr Gly Glu Lys Leu Ser Glu Thr Glu Ala Ala  
 420 425 430  
 Asp Ser Lys Asn Leu Thr Ser Lys Leu Leu Gln Pro Val Thr Leu Ser  
 435 440 445  
 Gly Gly Thr Leu Ser Leu Lys His Gly Val Thr Leu Gln Thr Gln Ala  
 450 455 460  
 Phe Thr Gln Gln Ala Asp Ser Arg Leu Glu Met Asp Val Gly Thr Thr  
 465 470 475 480  
 Leu Glu Pro Ala Asp Thr Ser Thr Ile Asn Asn Leu Val Ile Asn Ile  
 485 490 495  
 Ser Ser Ile Asp Gly Ala Lys Lys Ala Lys Ile Glu Thr Lys Ala Thr  
 500 505 510  
 Ser Lys Asn Leu Thr Leu Ser Gly Thr Ile Thr Leu Leu Asp Pro Thr  
 515 520 525  
 Gly Thr Phe Tyr Glu Asn His Ser Leu Arg Asn Pro Gln Ser Tyr Asp  
 530 535 540  
 Ile Leu Glu Leu Lys Ala Ser Gly Thr Val Thr Ser Thr Ala Val Thr  
 545 550 555 560  
 Pro Asp Pro Ile Met Gly Glu Lys Phe His Tyr Gly Tyr Gln Gly Thr  
 565 570 575  
 Trp Gly Pro Ile Val Trp Gly Thr Gly Ala Ser Thr Thr Ala Thr Phe  
 580 585 590  
 Asn Trp Thr Lys Thr Gly Tyr Ile Pro Asn Pro Glu Arg Ile Gly Ser  
 595 600 605  
 Leu Val Pro Asn Ser Leu Trp Asn Ala Phe Ile Asp Ile Ser Ser Leu  
 610 615 620  
 His Tyr Leu Met Glu Thr Ala Asn Glu Gly Leu Gln Gly Asp Arg Ala  
 625 630 635 640  
 Phe Trp Cys Ala Gly Leu Ser Asn Phe Phe His Lys Asp Ser Thr Lys  
 645 650 655  
 Thr Arg Arg Gly Phe Arg His Leu Ser Gly Gly Tyr Val Ile Gly Gly  
 660 665 670  
 Asn Leu His Thr Cys Ser Asp Lys Ile Leu Ser Ala Ala Phe Cys Gln  
 675 680 685  
 Leu Phe Gly Arg Asp Arg Asp Tyr Phe Val Ala Lys Asn Gln Gly Thr  
 690 695 700

Val Tyr Gly Gly Thr Leu Tyr Tyr Gln His Asn Glu Thr Tyr Ile Ser  
705 710 715 720  
Leu Pro Cys Lys Leu Arg Pro Cys Ser Leu Ser Tyr Val Pro Thr Glu  
725 730 735  
Ile Pro Val Leu Phe Ser Gly Asn Leu Ser Tyr Thr His Thr Asp Asn  
740 745 750  
Asp Leu Lys Thr Lys Tyr Thr Thr Tyr Pro Thr Val Lys Gly Ser Trp  
755 760 765  
Gly Asn Asp Ser Phe Ala Leu Glu Phe Gly Gly Arg Ala Pro Ile Cys  
770 775 780  
Leu Asp Glu Ser Ala Leu Phe Glu Gln Tyr Met Pro Phe Met Lys Leu  
785 790 795 800  
Gln Phe Val Tyr Ala His Gln Glu Gly Phe Lys Glu Gln Gly Thr Glu  
805 810 815  
Ala Arg Glu Phe Gly Ser Ser Arg Leu Val Asn Leu Ala Leu Pro Ile  
820 825 830  
Gly Ile Arg Phe Asp Lys Glu Ser Asp Cys Gln Asp Ala Thr Tyr Asn  
835 840 845  
Leu Thr Leu Gly Tyr Thr Val Asp Leu Val Arg Ser Asn Pro Asp Cys  
850 855 860  
Thr Thr Thr Leu Arg Ile Ser Gly Asp Ser Trp Lys Thr Phe Gly Thr  
865 870 875 880  
Asn Leu Ala Arg Gln Ala Leu Val Leu Arg Ala Gly Asn His Phe Cys  
885 890 895  
Phe Asn Ser Asn Phe Glu Ala Phe Ser Gln Phe Ser Phe Glu Leu Arg  
900 905 910  
Gly Ser Ser Arg Asn Tyr Asn Val Asp Leu Gly Ala Lys Tyr Gln Phe  
915 920 925

## (2) INFORMATION FOR SEQ ID NO:11:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2757 base pairs  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

ATGAGATCGT	CTTTTTCCTT	GTTATTAATA	TCTTCATCTC	TAGCCTTTCC	TCTCTTAATG	60
AGTGTTCCTG	CAGATGCTGC	CGATCTCACA	TTAGGGAGTC	GTGACAGTTA	TAATGGTGAT	120
ACAAGCACCA	CAGAATTTAC	TCCTAAAGCG	GCAACTTCTG	ATGCTAGTGG	CACGACCTAT	180
ATTCTCGATG	GGGATGTCTC	GATAAGCCAA	GCAGGGAAAC	AAACGAGCTT	AACCACAAGT	240
TGTTTTTCTA	ACACTGCAGG	AAATCTTACC	TTCTTAGGGA	ACGGATTTTC	TCTTCATTTT	300
GACAATATTA	TTTCGTCTAC	TGTTGCAGGT	GTTGTTGTGA	GCAATACAGC	AGCTTCTGGG	360
ATTACGAAAT	TCTCAGGATT	TTCAACTCTT	CGGATGCTTG	CAGCTCCTAG	GACCACAGGT	420
AAAGGAGCCA	TTAAAATTAC	CGATGGTCTG	GTGTTTGAGA	GTATAGGGAA	TCTTGACCAA	480
AATGAAAATG	CCTCTAGTGA	AAATGGGGGA	GCCATCAATA	CGAAGACTTT	GTCTTTGACT	540
GGGAGTACGC	GGTTTGTAGC	GTTCTTGGC	AATAGCTCGT	CGCAACAAGG	GGGAGCGATC	600
TATGCTTCTG	GTGACTCTGT	GATTTCTGAG	AATGCAGGAA	TCTTGAGCTT	CGGAAACAAC	660
AGTGCGACAA	CATCAGGAGG	CGCGATCTCT	GCTGAAGGGA	ACCTTGTGAT	CTCCAATAAC	720
CAAAATATCT	TTTTCGATGG	CTGCAAAGCA	ACTACAAATG	GCGGAGCTAT	TGATTGTAAAC	780
AAAGCAGGGG	CGAACCCAGA	CCCTATCTTG	ACTCTTTCAG	GAAATGAGAG	CCTGCATTTT	840
CTGAATAACA	CAGCAGGAAA	TAGTGGAGGT	GCGATTATATA	CCAAAAAATT	GGTGTATATCC	900
TCAGGACGAG	GAGGAGTGTT	ATTTTCTAAC	AACAAAGCTG	CGAATGCTAC	TCCTAAAGGA	960

GGGGCAATTG CGATTCTAGA TTCTGGAGAG ATTAGCATTT CTGCAGATCT CGGCAATATC 1020  
 ATTTTCGAGG GCAATACTAC GAGCACTACA GGAAGTCCTG CGAGTGTGAC CAGAAATGCT 1080  
 ATAGATCTTG CATCGAATGC AAAATTTTAA AATCTCCGAG CGACTCGGGG AAATAAAGTT 1140  
 ATTTTCTATG ATCCTATCAC GAGCTCAGGA GCTACTGATA AGCTCTCTTT GAATAAAGCT 1200  
 GACGCAGGAT CTGGAAATAC CTATGAAGGC TACATCGTTT TCTCTGGAGA GAAACTCTCA 1260  
 GAAGAGGAAC TTAAGAAACC TGACAATCTG AAGTCTACAT TTACACAGGC TGTAGAGCTT 1320  
 GCTGCAGGTG CCTTAGTATT GAAAGATGGA GTGACTGTAG TTGCAAATAC TATAACGCAG 1380  
 GTCGAGGGAT CGAAAGTCGT TATGGATGGA GGGACTACTT TTGAGGCAAG CGCTGAGGGG 1440  
 GTCACTCTCA ATGGCCTAGC CATTAATATA GATTCCTTAG ATGGGACAAA TAAAGCTATC 1500  
 ATTAAGGCGA CGGCAGCAAG TAAGGATGTT GCCTTATCAG GGCCTATCAT GCTTGTAGAT 1560  
 GCTCAGGGGA ACTATTATGA GCATCATAAT CTCAGTCAAC AGCAGGTCTT TCCTTTAATA 1620  
 GAGCTTTCTG CACAAGGAAC GATGACTACT ACAGATATCC CCGATACCCC AATTCTAAAT 1680  
 ACTACGAATC ACTATGGGTA TCAAGGAACT GGAATAATTG TTTGGGTCGA CGATGCAACT 1740  
 GCAAAAACAA AAAATGCTAC CTTAACTTGG ACTAAAACAG GATACAAGCC GAATCCAGAA 1800  
 CGTCAGGGAC CTTTGGTTCC TAATAGCCTG TGGGGTTCTT TTGTCGATGT CCGCTCCATT 1860  
 CAGAGCCTCA TGGACCGGAG CACAAGTTCG TTATCTTCGT CAACAAATTT GTGGGTATCA 1920  
 GGAATCGGGG ACTTTTGTGA TGAAGATCAG AAAGGAAACC AACGTAGTTA TCGTCATTCT 1980  
 AGCGCGGGTT ATGCATTAGG AGGAGGATTC TTCACGGCTT CTGAAAATTT CTTTAAATTT 2040  
 GCTTTTTGTC AGCTTTTTTG CTACGACAAG GACCATCTTG TGGCTAAGAA CCATACCCAT 2100  
 GTATATGCAG GGGCAATGAG TTACCGACAC CTCGGAGAGT CTAAGACCCT CGCTAAGATT 2160  
 TTGTCAGGAA ATTCTGACTC CCTACCTTTT GTCTTCAATG CTCGGTTTGC TTATGGCCAT 2220  
 ACCGACAATA ACATGACCAC AAAGTACACT GGCTATTCTC CTGTTAAGGG AAGCTGGGGA 2280  
 AATGATGCCT TCGGTATAGA ATGTGGAGGA GCTATCCCGG TAGTTGCTTC AGGACGTCGG 2340  
 TCTTGGGTGG ATACCCACAC GCCATTCTA AACCTAGAGA TGATCTATGC ACATCAGAAT 2400  
 GACTTTAAGG AAAACGGCAC AGAAGGCCGT TCTTTCCAAA GTGAAGACCT CTTCAATCTA 2460  
 GCGGTTCTCTG TAGGGATAAA ATTTGAGAAA TTCTCCGATA AGTCTACGTA TGATCTCTCC 2520  
 ATAGCTTACG TTCCCGATGT GATTCGTAAT GATCCAGGCT GCACGACAAC TCTTATGGTT 2580  
 TCTGGGGATT CTGGTTCGAC ATGTGGTACA AGCTTGCTTA GACAAGCTCT TCTTGTACGT 2640  
 GCTGGAAATC ATCATGCCTT TGCTTCAAAC TTTGAAGTTT TCAGTCAGTT TGAAGTCGAG 2700  
 TTGCGAGGTT CTTCTCGTAG CTATGCTATC GATCTTGGAG GAAGATTCGG ATTTTAA 2757

## (2) INFORMATION FOR SEQ ID NO:12:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 918 amino acids  
 (B) TYPE: amino acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:

Met Arg Ser Ser Phe Ser Leu Leu Leu Ile Ser Ser Ser Leu Ala Phe  
 1 5 10 15  
 Pro Leu Leu Met Ser Val Ser Ala Asp Ala Ala Asp Leu Thr Leu Gly  
 20 25 30  
 Ser Arg Asp Ser Tyr Asn Gly Asp Thr Thr Thr Thr Glu Phe Thr Pro  
 35 40 45  
 Lys Ala Ala Thr Ser Asp Ala Ser Gly Thr Tyr Ile Leu Asp Gly  
 50 55 60  
 Asp Val Ser Ile Ser Gln Ala Gly Lys Gln Thr Ser Leu Thr Thr Ser  
 65 70 75 80  
 Cys Phe Ser Asn Thr Ala Gly Asn Leu Thr Phe Leu Gly Asn Gly Phe  
 85 90 95  
 Ser Leu His Phe Asp Asn Ile Ile Ser Ser Thr Val Ala Gly Val Val  
 100 105 110

Val Ser Asn Thr Ala Ala Ser Gly Ile Thr Lys Phe Ser Gly Phe Ser  
 115 120 125  
 Thr Leu Arg Met Leu Ala Ala Pro Arg Thr Thr Gly Lys Gly Ala Ile  
 130 135 140  
 Lys Ile Thr Asp Gly Leu Val Phe Glu Ser Ile Gly Asn Leu Asp Gln  
 145 150 155 160  
 Asn Glu Asn Ala Ser Ser Glu Asn Gly Gly Ala Ile Asn Thr Lys Thr  
 165 170 175  
 Leu Ser Leu Thr Gly Ser Thr Arg Phe Val Ala Phe Leu Gly Asn Ser  
 180 185 190  
 Ser Ser Gln Gln Gly Gly Ala Ile Tyr Ala Ser Gly Asp Ser Val Ile  
 195 200 205  
 Ser Glu Asn Ala Gly Ile Leu Ser Phe Gly Asn Asn Ser Ala Thr Thr  
 210 215 220  
 Ser Gly Gly Ala Ile Ser Ala Glu Gly Asn Leu Val Ile Ser Asn Asn  
 225 230 235 240  
 Gln Asn Ile Phe Phe Asp Gly Cys Lys Ala Thr Thr Asn Gly Gly Ala  
 245 250 255  
 Ile Asp Cys Asn Lys Ala Gly Ala Asn Pro Asp Pro Ile Leu Thr Leu  
 260 265 270  
 Ser Gly Asn Glu Ser Leu His Phe Leu Asn Asn Thr Ala Gly Asn Ser  
 275 280 285  
 Gly Gly Ala Ile Tyr Thr Lys Lys Leu Val Leu Ser Ser Gly Arg Gly  
 290 295 300  
 Gly Val Leu Phe Ser Asn Asn Lys Ala Ala Asn Ala Thr Pro Lys Gly  
 305 310 315 320  
 Gly Ala Ile Ala Ile Leu Asp Ser Gly Glu Ile Ser Ile Ser Ala Asp  
 325 330 335  
 Leu Gly Asn Ile Ile Phe Glu Gly Asn Thr Thr Ser Thr Thr Gly Ser  
 340 345 350  
 Pro Ala Ser Val Thr Arg Asn Ala Ile Asp Leu Ala Ser Asn Ala Lys  
 355 360 365  
 Phe Leu Asn Leu Arg Ala Thr Arg Gly Asn Lys Val Ile Phe Tyr Asp  
 370 375 380  
 Pro Ile Thr Ser Ser Gly Ala Thr Asp Lys Leu Ser Leu Asn Lys Ala  
 385 390 395 400  
 Asp Ala Gly Ser Gly Asn Thr Tyr Glu Gly Tyr Ile Val Phe Ser Gly  
 405 410 415  
 Glu Lys Leu Ser Glu Glu Glu Leu Lys Lys Pro Asp Asn Leu Lys Ser  
 420 425 430  
 Thr Phe Thr Gln Ala Val Glu Leu Ala Ala Gly Ala Leu Val Leu Lys  
 435 440 445  
 Asp Gly Val Thr Val Val Ala Asn Thr Ile Thr Gln Val Glu Gly Ser  
 450 455 460  
 Lys Val Val Met Asp Gly Gly Thr Thr Phe Glu Ala Ser Ala Glu Gly  
 465 470 475 480  
 Val Thr Leu Asn Gly Leu Ala Ile Asn Ile Asp Ser Leu Asp Gly Thr  
 485 490 495  
 Asn Lys Ala Ile Ile Lys Ala Thr Ala Ala Ser Lys Asp Val Ala Leu  
 500 505 510  
 Ser Gly Pro Ile Met Leu Val Asp Ala Gln Gly Asn Tyr Tyr Glu His  
 515 520 525  
 His Asn Leu Ser Gln Gln Gln Val Phe Pro Leu Ile Glu Leu Ser Ala  
 530 535 540  
 Gln Gly Thr Met Thr Thr Thr Asp Ile Pro Asp Thr Pro Ile Leu Asn  
 545 550 555 560  
 Thr Thr Asn His Tyr Gly Tyr Gln Gly Thr Gly Ile Ile Val Trp Val

565 570 575  
 Asp Asp Ala Thr Ala Lys Thr Lys Asn Ala Thr Leu Thr Trp Thr Lys  
 580 585 590  
 Thr Gly Tyr Lys Pro Asn Pro Glu Arg Gln Gly Pro Leu Val Pro Asn  
 595 600 605  
 Ser Leu Trp Gly Ser Phe Val Asp Val Arg Ser Ile Gln Ser Leu Met  
 610 615 620  
 Asp Arg Ser Thr Ser Ser Leu Ser Ser Ser Thr Asn Leu Trp Val Ser  
 625 630 635 640  
 Gly Ile Ala Asp Phe Leu His Glu Asp Gln Lys Gly Asn Gln Arg Ser  
 645 650 655  
 Tyr Arg His Ser Ser Ala Gly Tyr Ala Leu Gly Gly Gly Phe Phe Thr  
 660 665 670  
 Ala Ser Glu Asn Phe Phe Asn Phe Ala Phe Cys Gln Leu Phe Gly Tyr  
 675 680 685  
 Asp Lys Asp His Leu Val Ala Lys Asn His Thr His Val Tyr Ala Gly  
 690 695 700  
 Ala Met Ser Tyr Arg His Leu Gly Glu Ser Lys Thr Leu Ala Lys Ile  
 705 710 715 720  
 Leu Ser Gly Asn Ser Asp Ser Leu Pro Phe Val Phe Asn Ala Arg Phe  
 725 730 735  
 Ala Tyr Gly His Thr Asp Asn Asn Met Thr Thr Lys Tyr Thr Gly Tyr  
 740 745 750  
 Ser Pro Val Lys Gly Ser Trp Gly Asn Asp Ala Phe Gly Ile Glu Cys  
 755 760 765  
 Gly Gly Ala Ile Pro Val Val Ala Ser Gly Arg Arg Ser Trp Val Asp  
 770 775 780  
 Thr His Thr Pro Phe Leu Asn Leu Glu Met Ile Tyr Ala His Gln Asn  
 785 790 795 800  
 Asp Phe Lys Glu Asn Gly Thr Glu Gly Arg Ser Phe Gln Ser Glu Asp  
 805 810 815  
 Leu Phe Asn Leu Ala Val Pro Val Gly Ile Lys Phe Glu Lys Phe Ser  
 820 825 830  
 Asp Lys Ser Thr Tyr Asp Leu Ser Ile Ala Tyr Val Pro Asp Val Ile  
 835 840 845  
 Arg Asn Asp Pro Gly Cys Thr Thr Thr Leu Met Val Ser Gly Asp Ser  
 850 855 860  
 Trp Ser Thr Cys Gly Thr Ser Leu Ser Arg Gln Ala Leu Leu Val Arg  
 865 870 875 880  
 Ala Gly Asn His His Ala Phe Ala Ser Asn Phe Glu Val Phe Ser Gln  
 885 890 895  
 Phe Glu Val Glu Leu Arg Gly Ser Ser Arg Ser Tyr Ala Ile Asp Leu  
 900 905 910  
 Gly Gly Arg Phe Gly Phe  
 915

## (2) INFORMATION FOR SEQ ID NO:13:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2787 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:

ATGAAATCCT	CTCTTCATTG	GTTTGTAATC	TCGTCACTCT	TAGCACTTCC	CTTGTCACTA	60
AATTTCTCTG	CGTTTGCTGC	TGTTGTTGAA	ATCAATCTAG	GACCTACCAA	TAGCTTCTCT	120
GGACCAGGAA	CCTACACTCC	TCCAGCCCAA	ACAACAAATG	CAGATGGAAC	TATCTATAAT	180
CTAACAGGGG	ATGTCTCAAT	CACCAATGCA	GGATCTCCGA	CAGCTCTAAC	CGCTTCCTGC	240
TTTAAAGAAA	CTACTGGGAA	TCTTTCTTTC	CAAGGCCACG	GCTACCAATT	TCTCCTACAA	300
AATATCGATG	CGGGAGCGAA	CTGTACCTTT	ACCAATACAG	CTGCAAATAA	GCTTCTCTCC	360
TTTTCAAGAT	TCTCCTATTT	GTCACTAATA	CAAACCACGA	ATGCTACCAC	AGGAACAGGA	420
GCCATCAAGT	CCACAGGAGC	TTGTTCTATT	CAGTCGAACT	ATAGTTGCTA	CTTTGGCCAA	480
AACTTTTCTA	ATGACAATGG	AGGCGCCCTC	CAAGGCAGCT	CTATCAGTCT	ATCGCTAAAC	540
CCCAACCTAA	CGTTTGCCAA	AAACAAAGCA	ACGCAAAAAG	GGGGTGCCCT	CTATTCCACG	600
GGAGGGATTA	CAATTAACAA	TACGTTAAAC	TCAGCATCAT	TTTCTGAAA	TACCGCGGCG	660
AACAATGGCG	GAGCCATTTA	CACGGAAGCT	AGCAGTTTTA	TTAGCAGCAA	CAAAGCAATT	720
AGCTTTATAA	ACAATAGTGT	GACCGCAACC	TCAGCTACAG	GGGGAGCCAT	TTACTGTAGT	780
AGTACATCAG	CCCCCAAACC	AGTCTTAACT	CTATCAGACA	ACGGGGAAC	GAACTTTATA	840
GGAAATACAG	CAATTACTAG	TGGTGGGGCG	ATTTATACTG	ACAATCTAGT	TCTTTCTTCT	900
GGAGGACCTA	CGCTTTTTTA	AAACAACCTCT	GCTATAGATA	CTGCAGCTCC	CTTAGGAGGA	960
GCAATTGCGA	TTGCTGACTC	TGGATCTTTG	AGTCTTTCGG	CTCTTGGTGG	AGACATCACT	1020
TTTGAAGGAA	ACACAGTAGT	CAAAGGAGCT	TCTTCGAGTC	AGACCACTAC	CAGAAATTCT	1080
ATTAACATCG	GAAACACCAA	TGCTAAGATT	GTACAGCTGC	GAGCCTCTCA	AGGCAATACT	1140
ATCTACTTCT	ATGATCCTAT	AACAATAAC	CATACTGCAG	CTCTCTCAGA	TGCTCTAAAC	1200
TTAAATGGTC	CTGACCTTGC	AGGGAATCCT	GCATATCAAG	GAACCATCGT	ATTTTCTGGA	1260
GAGAAGCTCT	CGGAAGCAGA	AGCTGCAGAA	GCTGATAATC	TCAAATCTAC	AATTCAGCAA	1320
CCTCTAACTC	TTGCGGGAGG	GCAACTCTCT	CTTAAATCAG	GAGTCACTCT	AGTTGCTAAG	1380
TCCTTTTCGC	AATCTCCGGG	CTCTACCCTC	CTCATGGATG	CAGGGACCAC	ATTAGAAACC	1440
GCTGTGGGA	TCACTTCAAA	TAATCTTGT	CTCAATGTAG	ATTCTTAAA	AGAGACCAAG	1500
AAGGCTACGC	TAAAGCAAC	ACAAGCAAGT	CAGACAGTCA	CTTTATCTGG	ATCGCTCTCT	1560
CTTGTAGATC	CTTCTGAAA	TGTCTACGAA	GATGTCTCTT	GGAATAACCC	TCAAGTCTTT	1620
TCTTGTCTCA	CTCTTACTGC	TGACGACCCC	GCGAATATTC	ACATCACAGA	CTTAGCTGCT	1680
GATCCCCTAG	AAAAAATCC	TATCCATTGG	GGATACCAAG	GGAATTGGGC	ATTATCTTGG	1740
CAAGAGGATA	CTGCGACTAA	ATCCAAAGCA	GCGACTCTTA	CCTGGACAAA	AACAGGATAC	1800
AATCCGAATC	CTGAGCGTCG	TGGAACCTTA	GTTGCTAACA	CGCTATGGGG	ATCCTTTGTT	1860
GATGTGCGCT	CCATACAACA	GCTTGTAGCC	ACTAAAGTAC	GCCAATCTCA	AGAAACTCGC	1920
GGCATCTGGT	GTGAAGGGAT	CTCGAAGTTC	TTCCATAAAG	ATAGCACGAA	GATAAATAAA	1980
GGTTTTCGCC	ACATAAGTGC	AGGTTATGTT	GTAGGAGCGA	CTACAACATT	AGCTTCTGAT	2040
AATCTTATCA	CTGCAGCCTT	CTGCCAATTA	TTCCGGGAAAG	ATAGAGATCA	CTTTATAAAT	2100
AAAAATAGAG	CTTCTGCCTA	TGCAGCTTCT	CTCCATCTCC	AGCATCTAGC	GACCTTGTCT	2160
TCTCCAAGCT	TGTTACGCTA	CCTTCCTGGA	TCTGAAAGTG	AGCAGCCTGT	CCTCTTTGAT	2220
GCTCAGATCA	GCTATATCTA	TAGTAAAAAT	ACTATGAAAA	CCTATTACAC	CCAAGCACCA	2280
AAGGGAGAGA	GCTCGTGGTA	TAATGACGGT	TGCGCTCTGG	AACTTGCGAG	CTCCCTACCA	2340
CACACTGCTT	TAAGCCATGA	GGGTCTCTTC	CACGCGTATT	TTCTTTTCAT	CAAAGTAGAA	2400
GCTTCGTACA	TACACCAAGA	TAGCTTCAAA	GAACGTAATA	CTACCTTGGT	ACGATCTTTC	2460
GATAGCGGTG	ATTTAATTAA	CGTCTCTGTG	CCTATTGGAA	TTACCTTCGA	GAGATTCTCG	2520
AGAAACGAGC	GTGCGTCTTA	CGAAGCTACT	GTCATCTACG	TTGCCGATGT	CTATCGTAAG	2580
AATCCTGACT	GCACGACAGC	TCTCCTAATC	AACAATACCT	CGTGGAAAAC	TACAGGAACG	2640
AATCTCTCAA	GACAAGCTGG	TATCGGAAGA	GCAGGGATCT	TTTATGCCTT	CTCTCCAAAT	2700
CTTGAGGTCA	CAAGTAACCT	ATCTATGGAA	ATTCGTGGAT	CTTCACGCAG	CTACAATGCA	2760
GATCTTGGAG	GTAAGTTCCA	GTTCTAA				2787

## (2) INFORMATION FOR SEQ ID NO:14:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 928 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide



(xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:

```

Met Lys Ser Ser Leu His Trp Phe Val Ile Ser Ser Ser Leu Ala Leu
 1          5          10          15
Pro Leu Ser Leu Asn Phe Ser Ala Phe Ala Ala Val Val Glu Ile Asn
          20          25          30
Leu Gly Pro Thr Asn Ser Phe Ser Gly Pro Gly Thr Tyr Thr Pro Pro
          35          40          45
Ala Gln Thr Thr Asn Ala Asp Gly Thr Ile Tyr Asn Leu Thr Gly Asp
          50          55          60
Val Ser Ile Thr Asn Ala Gly Ser Pro Thr Ala Leu Thr Ala Ser Cys
65          70          75          80
Phe Lys Glu Thr Thr Gly Asn Leu Ser Phe Gln Gly His Gly Tyr Gln
          85          90          95
Phe Leu Leu Gln Asn Ile Asp Ala Gly Ala Asn Cys Thr Phe Thr Asn
          100          105          110
Thr Ala Ala Asn Lys Leu Leu Ser Phe Ser Gly Phe Ser Tyr Leu Ser
          115          120          125
Leu Ile Gln Thr Thr Asn Ala Thr Thr Gly Thr Gly Ala Ile Lys Ser
          130          135          140
Thr Gly Ala Cys Ser Ile Gln Ser Asn Tyr Ser Cys Tyr Phe Gly Gln
145          150          155          160
Asn Phe Ser Asn Asp Asn Gly Gly Ala Leu Gln Gly Ser Ser Ile Ser
          165          170          175
Leu Ser Leu Asn Pro Asn Leu Thr Phe Ala Lys Asn Lys Ala Thr Gln
          180          185          190
Lys Gly Gly Ala Leu Tyr Ser Thr Gly Gly Ile Thr Ile Asn Asn Thr
          195          200          205
Leu Asn Ser Ala Ser Phe Ser Glu Asn Thr Ala Ala Asn Asn Gly Gly
210          215          220
Ala Ile Tyr Thr Glu Ala Ser Ser Phe Ile Ser Ser Asn Lys Ala Ile
225          230          235          240
Ser Phe Ile Asn Asn Ser Val Thr Ala Thr Ser Ala Thr Gly Gly Ala
          245          250          255
Ile Tyr Cys Ser Ser Thr Ser Ala Pro Lys Pro Val Leu Thr Leu Ser
260          265          270
Asp Asn Gly Glu Leu Asn Phe Ile Gly Asn Thr Ala Ile Thr Ser Gly
275          280          285
Gly Ala Ile Tyr Thr Asp Asn Leu Val Leu Ser Ser Gly Gly Pro Thr
290          295          300
Leu Phe Lys Asn Asn Ser Ala Ile Asp Thr Ala Ala Pro Leu Gly Gly
305          310          315          320
Ala Ile Ala Ile Ala Asp Ser Gly Ser Leu Ser Leu Ser Ala Leu Gly
          325          330          335
Gly Asp Ile Thr Phe Glu Gly Asn Thr Val Val Lys Gly Ala Ser Ser
          340          345          350
Ser Gln Thr Thr Thr Arg Asn Ser Ile Asn Ile Gly Asn Thr Asn Ala
          355          360          365
Lys Ile Val Gln Leu Arg Ala Ser Gln Gly Asn Thr Ile Tyr Phe Tyr
          370          375          380
Asp Pro Ile Thr Thr Asn His Thr Ala Ala Leu Ser Asp Ala Leu Asn
385          390          395          400
Leu Asn Gly Pro Asp Leu Ala Gly Asn Pro Ala Tyr Gln Gly Thr Ile
          405          410          415
Val Phe Ser Gly Glu Lys Leu Ser Glu Ala Glu Ala Ala Glu Ala Asp
          420          425          430
Asn Leu Lys Ser Thr Ile Gln Gln Pro Leu Thr Leu Ala Gly Gly Gln

```

435 440 445  
 Leu Ser Leu Lys Ser Gly Val Thr Leu Val Ala Lys Ser Phe Ser Gln  
 450 455 460  
 Ser Pro Gly Ser Thr Leu Leu Met Asp Ala Gly Thr Thr Leu Glu Thr  
 465 470 475 480  
 Ala Asp Gly Ile Thr Ile Asn Asn Leu Val Leu Asn Val Asp Ser Leu  
 485 490 495  
 Lys Glu Thr Lys Lys Ala Thr Leu Lys Ala Thr Gln Ala Ser Gln Thr  
 500 505 510  
 Val Thr Leu Ser Gly Ser Leu Ser Leu Val Asp Pro Ser Gly Asn Val  
 515 520 525  
 Tyr Glu Asp Val Ser Trp Asn Asn Pro Gln Val Phe Ser Cys Leu Thr  
 530 535 540  
 Leu Thr Ala Asp Asp Pro Ala Asn Ile His Ile Thr Asp Leu Ala Ala  
 545 550 555 560  
 Asp Pro Leu Glu Lys Asn Pro Ile His Trp Gly Tyr Gln Gly Asn Trp  
 565 570 575  
 Ala Leu Ser Trp Gln Glu Asp Thr Ala Thr Lys Ser Lys Ala Ala Thr  
 580 585 590  
 Leu Thr Trp Thr Lys Thr Gly Tyr Asn Pro Asn Pro Glu Arg Arg Gly  
 595 600 605  
 Thr Leu Val Ala Asn Thr Leu Trp Gly Ser Phe Val Asp Val Arg Ser  
 610 615 620  
 Ile Gln Gln Leu Val Ala Thr Lys Val Arg Gln Ser Gln Glu Thr Arg  
 625 630 635 640  
 Gly Ile Trp Cys Glu Gly Ile Ser Asn Phe Phe His Lys Asp Ser Thr  
 645 650 655  
 Lys Ile Asn Lys Gly Phe Arg His Ile Ser Ala Gly Tyr Val Val Gly  
 660 665 670  
 Ala Thr Thr Thr Leu Ala Ser Asp Asn Leu Ile Thr Ala Ala Phe Cys  
 675 680 685  
 Gln Leu Phe Gly Lys Asp Arg Asp His Phe Ile Asn Lys Asn Arg Ala  
 690 695 700  
 Ser Ala Tyr Ala Ala Ser Leu His Leu Gln His Leu Ala Thr Leu Ser  
 705 710 715 720  
 Ser Pro Ser Leu Leu Arg Tyr Leu Pro Gly Ser Glu Ser Glu Gln Pro  
 725 730 735  
 Val Leu Phe Asp Ala Gln Ile Ser Tyr Ile Tyr Ser Lys Asn Thr Met  
 740 745 750  
 Lys Thr Tyr Tyr Thr Gln Ala Pro Lys Gly Glu Ser Ser Trp Tyr Asn  
 755 760 765  
 Asp Gly Cys Ala Leu Glu Leu Ala Ser Ser Leu Pro His Thr Ala Leu  
 770 775 780  
 Ser His Glu Gly Leu Phe His Ala Tyr Phe Pro Phe Ile Lys Val Glu  
 785 790 795 800  
 Ala Ser Tyr Ile His Gln Asp Ser Phe Lys Glu Arg Asn Thr Thr Leu  
 805 810 815  
 Val Arg Ser Phe Asp Ser Gly Asp Leu Ile Asn Val Ser Val Pro Ile  
 820 825 830  
 Gly Ile Thr Phe Glu Arg Phe Ser Arg Asn Glu Arg Ala Ser Tyr Glu  
 835 840 845  
 Ala Thr Val Ile Tyr Val Ala Asp Val Tyr Arg Lys Asn Pro Asp Cys  
 850 855 860  
 Thr Thr Ala Leu Leu Ile Asn Asn Thr Ser Trp Lys Thr Thr Gly Thr  
 865 870 875 880  
 Asn Leu Ser Arg Gln Ala Gly Ile Gly Arg Ala Gly Ile Phe Tyr Ala  
 885 890 895

Phe Ser Pro Asn Leu Glu Val Thr Ser Asn Leu Ser Met Glu Ile Arg  
 900 905 910  
 Gly Ser Ser Arg Ser Tyr Asn Ala Asp Leu Gly Gly Lys Phe Gln Phe  
 915 920 925

## (2) INFORMATION FOR SEQ ID NO:15:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2793 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:

ATGAAAATAC	CCTGACAA	ACTCCTGATC	TCTTCGACTC	TTGTCACTCC	CATTCTATTG	60
AGCATTGCAA	CTTACGGAGC	AGATGCTTCT	TTATCCCCTA	CAGATAGCTT	TGATGGAGCG	120
GGCGGCTCTA	CATTACTCC	AAAATCTACA	GCAGATGCCA	ATGGAACGAA	CTATGTCTTA	180
TCAGGAAATG	TCTATATAAA	CGATGCTGGG	AAAGGCACAG	CATTAAACAGG	CTGCTGCTTT	240
ACAGAAACTA	CGGGTGATCT	GACATTTACT	GGAAAGGGAT	ACTCATTTTC	ATTCAACACG	300
GTAGATGCGG	GTTCAATGC	AGGAGCTGCG	GCAAGCACAA	CTGCTGATAA	AGCCCTAACA	360
TTCACAGGAT	TTTCTAACCT	TTCTTCATT	GCAGCTCCTG	GAACACAGT	TGCTTCAGGA	420
AAAAGTACTT	TAAGTCTGTC	AGGAGCCTTA	AATCTTACCG	ATAATGGAAC	GATTCTCTTT	480
AGCCAAAACG	TCTCCAATGA	AGCTAATAAC	AATGGCGGAG	CGATCACCAC	AAAAACTCTT	540
TCTATTTCTG	GGAATACCTC	TTCTATAACC	TTCACTAGTA	ATAGCGCAAA	AAAATTAGGT	600
GGAGCGATCT	ATAGCTCTGC	GGCTGCAAGT	ATTTCAGGAA	ACACCGGCCA	GTTAGTCTTT	660
ATGAATAATA	AAGGAGAAAC	TGGGGGCGGG	GCTCTGGGCT	TTGAAGCCAG	CTCCTCGATT	720
ACTCAAATA	GCTCCCTTTT	CTTCTCTGGA	AACACTGCAA	CAGATGCTGC	AGGCAAGGGC	780
GGGGCCATTT	ATTGTGAAAA	AACAGGAGAG	ACTCCTACTC	TTACTATCTC	TGGAAATAAA	840
AGTCTGACCT	TCGCCGAGAA	CTCTTCAGTA	ACTCAAGGCG	GAGCAATCTG	TGCCCATGGT	900
CTAGATCTTT	CCGCTGCTGG	CCCTACCCTA	TTTTCAAATA	ATAGATGCGG	GAACACAGCT	960
GCAGGCAAGG	GCGGCGCTAT	TGCAATTGCC	GACTCTGGAT	CTTTAAGTCT	CTCTGCAAT	1020
CAAGGAGACA	TCACGTTCCT	TGGCAACACT	CTAACCTCAA	CCTCCGCGCC	AACATCGACA	1080
CGGAATGCTA	TCTACCTGGG	ATCGTCAGCA	AAAATTACGA	ACTTAAGGGC	AGCCCAAGGC	1140
CAATCTATCT	ATTTCTATGA	TCCGATTGCA	TCTAACACCA	CAGGAGCTTC	AGACGTTCTG	1200
ACCATCAACC	AACCGGATAG	CAACTCGCCT	TTAGATTATT	CAGGAACGAT	TGTATTTTCT	1260
GGGGAAAAGC	TCTCTGCAGA	TGAAGCGAAA	GCTGCTGATA	ACTTCACATC	TATATTAAAG	1320
CAACCATTGG	CTCTAGCCTC	TGGAACCTTA	GCACTCAAAG	GAAATGTCGA	GTTAGATGTC	1380
AATGGTTTCA	CACAGACTGA	AGGCTCTACA	CTCCTCATGC	AACCAGGAAC	AAAGCTCAAA	1440
GCAGATACTG	AAGCTATCAG	TCTTACCAAA	CTTGTCGTTG	ATCTTTCTGC	CTTAGAGGGA	1500
AATAAGAGTG	TGTCCATTGA	AACAGCAGGA	GCCAACAAAA	CTATAACTCT	AACCTCTCCT	1560
CTTGTTTTCC	AAGATAGTAG	CGGCAATTTT	TATGAAAGCC	ATACGATAAA	CCAAGCCTTC	1620
ACGCAGCCTT	TGGTGGTATT	CACTGCTGCT	ACTGCTGCTA	GCGATATTTA	TATCGATGCG	1680
CTTCTCACTT	CTCCAGTACA	AACCTCAGAA	CCTCATTACG	GGTATCAGGG	ACATTGGGAA	1740
GCCACTTGGG	CAGACACATC	AACCTGCAAAA	TCAGGAACTA	TGACTTGGGT	AACCTACGGG	1800
TACAACCCTA	ATCCTGAGCG	TAGAGCTTCC	GTAGTTCCCG	ATTCATTATG	GGCATCCTTT	1860
ACTGACATTG	GCACTCTACA	GCAGATCATG	ACATCTCAAG	CGAATAGTAT	CTATCAGCAA	1920
CGAGGACTCT	GGGCATCAGG	AACCTGCGAAT	TTCTTCCATA	AGGATAAATC	AGGAACTAAC	1980
CAAGCATTCC	GACATAAAAG	CTACGGCTAT	ATTGTTGGAG	GAAGTGCTGA	AGATTTTCTT	2040
GAAAATATCT	TCAGTGTAGC	TTTCTGCCAG	CTCTTCGGTA	AAGATAAAGA	CCTGTTTATA	2100
GTTGAAAATA	CCTCTCATAA	CTATTTAGCG	TCGCTATACC	TGCAACATCG	AGCATTCTTA	2160
GGAGGACTTC	CCATGCCCTC	ATTTGGAAGT	ATCACCGACA	TGCTGAAAGA	TATTCCTCTC	2220
ATTTTGAATG	CCAGCTAAG	CTACAGCTAC	ACTAAAAATG	ATATGGATAC	TCGCTATACT	2280
TCCTATCCTG	AAGCTCAAGG	TTCTTGGACC	AATAATTCTG	GGGCTCTAGA	GCTCGGAGGA	2340
TCTCTGGCTC	TATATCTCCC	TAAAGAAGCA	CCGTTCTTCC	AGGGATATTT	CCCCTTCTTA	2400

AAGTTCCAGG	CAGTCTACAG	CCGCCAACAA	AACTTTAAAG	AGAGTGCGCG	TGAAGCCCGT	2460
GCTTTTGATG	ATGGAGACCT	AGTGAAGTGC	TCTATCCCTG	TCGGCATTGG	GTTAGAAAAA	2520
ATCTCCGAAG	ATGAAAAAAA	TAATTTTCGAG	ATTTCTCTAG	CCAACATTGG	TGATGTGTAT	2580
CGTAAAAATC	CCCGTTTCGCG	TACTTCTCTA	ATGGTCAGTG	GAGCCTCTTG	GACTTCGCTA	2640
TGTAAAAACC	TCGCACGACA	AGCCTTCTTA	GCAAGTGCTG	GAAGCCATCT	GACTCTCTCC	2700
CCTCATGTAG	AACTCTCTGG	GGAAGCTGCT	TATGAGCTTC	GTGGCTCAGC	ACACATCTAC	2760
AATGTAGATT	GTGGGCTAAG	ATACTCATTG	TAG			2793

## (2) INFORMATION FOR SEQ ID NO:16:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 930 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

Met	Lys	Ile	Pro	Leu	His	Lys	Leu	Leu	Ile	Ser	Ser	Thr	Leu	Val	Thr	1	5	10	15
Pro	Ile	Leu	Leu	Ser	Ile	Ala	Thr	Tyr	Gly	Ala	Asp	Ala	Ser	Leu	Ser	20	25	30	
Pro	Thr	Asp	Ser	Phe	Asp	Gly	Ala	Gly	Gly	Ser	Thr	Phe	Thr	Pro	Lys	35	40	45	
Ser	Thr	Ala	Asp	Ala	Asn	Gly	Thr	Asn	Tyr	Val	Leu	Ser	Gly	Asn	Val	50	55	60	
Tyr	Ile	Asn	Asp	Ala	Gly	Lys	Gly	Thr	Ala	Leu	Thr	Gly	Cys	Cys	Phe	65	70	75	80
Thr	Glu	Thr	Thr	Gly	Asp	Leu	Thr	Phe	Thr	Gly	Lys	Gly	Tyr	Ser	Phe	85	90	95	
Ser	Phe	Asn	Thr	Val	Asp	Ala	Gly	Ser	Asn	Ala	Gly	Ala	Ala	Ala	Ser	100	105	110	
Thr	Thr	Ala	Asp	Lys	Ala	Leu	Thr	Phe	Thr	Gly	Phe	Ser	Asn	Leu	Ser	115	120	125	
Phe	Ile	Ala	Ala	Pro	Gly	Thr	Val	Ala	Ser	Gly	Lys	Ser	Thr	Leu	130	135	140		
Ser	Ser	Ala	Gly	Ala	Leu	Asn	Leu	Thr	Asp	Asn	Gly	Thr	Ile	Leu	Phe	145	150	155	160
Ser	Gln	Asn	Val	Ser	Asn	Glu	Ala	Asn	Asn	Asn	Gly	Gly	Ala	Ile	Thr	165	170	175	
Thr	Lys	Thr	Leu	Ser	Ile	Ser	Gly	Asn	Thr	Ser	Ser	Ile	Thr	Phe	Thr	180	185	190	
Ser	Asn	Ser	Ala	Lys	Lys	Leu	Gly	Gly	Ala	Ile	Tyr	Ser	Ser	Ala	Ala	195	200	205	
Ala	Ser	Ile	Ser	Gly	Asn	Thr	Gly	Gln	Leu	Val	Phe	Met	Asn	Asn	Lys	210	215	220	
Gly	Glu	Thr	Gly	Gly	Gly	Ala	Leu	Gly	Phe	Glu	Ala	Ser	Ser	Ser	Ile	225	230	235	240
Thr	Gln	Asn	Ser	Ser	Leu	Phe	Phe	Ser	Gly	Asn	Thr	Ala	Thr	Asp	Ala	245	250	255	
Ala	Gly	Lys	Gly	Gly	Ala	Ile	Tyr	Cys	Glu	Lys	Thr	Gly	Glu	Thr	Pro	260	265	270	
Thr	Leu	Thr	Ile	Ser	Gly	Asn	Lys	Ser	Leu	Thr	Phe	Ala	Glu	Asn	Ser	275	280	285	
Ser	Val	Thr	Gln	Gly	Gly	Ala	Ile	Cys	Ala	His	Gly	Leu	Asp	Leu	Ser				

290	295	300
Ala Ala Gly Pro Thr Leu Phe Ser Asn Asn Arg Cys Gly Asn Thr Ala		
305	310	315
Ala Gly Lys Gly Gly Ala Ile Ala Ile Ala Asp Ser Gly Ser Leu Ser		
	325	330
Leu Ser Ala Asn Gln Gly Asp Ile Thr Phe Leu Gly Asn Thr Leu Thr		
	340	345
Ser Thr Ser Ala Pro Thr Ser Thr Arg Asn Ala Ile Tyr Leu Gly Ser		
	355	360
Ser Ala Lys Ile Thr Asn Leu Arg Ala Ala Gln Gly Gln Ser Ile Tyr		
	370	375
Phe Tyr Asp Pro Ile Ala Ser Asn Thr Thr Gly Ala Ser Asp Val Leu		
385	390	395
Thr Ile Asn Gln Pro Asp Ser Asn Ser Pro Leu Asp Tyr Ser Gly Thr		
	405	410
Ile Val Phe Ser Gly Glu Lys Leu Ser Ala Asp Glu Ala Lys Ala Ala		
	420	425
Asp Asn Phe Thr Ser Ile Leu Lys Gln Pro Leu Ala Leu Ala Ser Gly		
	435	440
Thr Leu Ala Leu Lys Gly Asn Val Glu Leu Asp Val Asn Gly Phe Thr		
	450	455
Gln Thr Glu Gly Ser Thr Leu Leu Met Gln Pro Gly Thr Lys Leu Lys		
465	470	475
Ala Asp Thr Glu Ala Ile Ser Leu Thr Lys Leu Val Val Asp Leu Ser		
	485	490
Ala Leu Glu Gly Asn Lys Ser Val Ser Ile Glu Thr Ala Gly Ala Asn		
	500	505
Lys Thr Ile Thr Leu Thr Ser Pro Leu Val Phe Gln Asp Ser Ser Gly		
	515	520
Asn Phe Tyr Glu Ser His Thr Ile Asn Gln Ala Phe Thr Gln Pro Leu		
	530	535
Val Val Phe Thr Ala Ala Thr Ala Ala Ser Asp Ile Tyr Ile Asp Ala		
545	550	555
Leu Leu Thr Ser Pro Val Gln Thr Pro Glu Pro His Tyr Gly Tyr Gln		
	565	570
Gly His Trp Glu Ala Thr Trp Ala Asp Thr Ser Thr Ala Lys Ser Gly		
	580	585
Thr Met Thr Trp Val Thr Thr Gly Tyr Asn Pro Asn Pro Glu Arg Arg		
	595	600
Ala Ser Val Val Pro Asp Ser Leu Trp Ala Ser Phe Thr Asp Ile Arg		
	610	615
Thr Leu Gln Gln Ile Met Thr Ser Gln Ala Asn Ser Ile Tyr Gln Gln		
625	630	635
Arg Gly Leu Trp Ala Ser Gly Thr Ala Asn Phe Phe His Lys Asp Lys		
	645	650
Ser Gly Thr Asn Gln Ala Phe Arg His Lys Ser Tyr Gly Tyr Ile Val		
	660	665
Gly Gly Ser Ala Glu Asp Phe Ser Glu Asn Ile Phe Ser Val Ala Phe		
	675	680
Cys Gln Leu Phe Gly Lys Asp Lys Asp Leu Phe Ile Val Glu Asn Thr		
	690	695
Ser His Asn Tyr Leu Ala Ser Leu Tyr Leu Gln His Arg Ala Phe Leu		
705	710	715
Gly Gly Leu Pro Met Pro Ser Phe Gly Ser Ile Thr Asp Met Leu Lys		
	725	730
Asp Ile Pro Leu Ile Leu Asn Ala Gln Leu Ser Tyr Ser Tyr Thr Lys		
	740	745
		750

Asn Asp Met Asp Thr Arg Tyr Thr Ser Tyr Pro Glu Ala Gln Gly Ser  
 755 760 765  
 Trp Thr Asn Asn Ser Gly Ala Leu Glu Leu Gly Gly Ser Leu Ala Leu  
 770 775 780  
 Tyr Leu Pro Lys Glu Ala Pro Phe Phe Gln Gly Tyr Phe Pro Phe Leu  
 785 790 795 800  
 Lys Phe Gln Ala Val Tyr Ser Arg Gln Gln Asn Phe Lys Glu Ser Gly  
 805 810 815  
 Ala Glu Ala Arg Ala Phe Asp Asp Gly Asp Leu Val Asn Cys Ser Ile  
 820 825 830  
 Pro Val Gly Ile Arg Leu Glu Lys Ile Ser Glu Asp Glu Lys Asn Asn  
 835 840 845  
 Phe Glu Ile Ser Leu Ala Asn Ile Gly Asp Val Tyr Arg Lys Asn Pro  
 850 855 860  
 Arg Ser Arg Thr Ser Leu Met Val Ser Gly Ala Ser Trp Thr Ser Leu  
 865 870 875 880  
 Cys Lys Asn Leu Ala Arg Gln Ala Phe Leu Ala Ser Ala Gly Ser His  
 885 890 895  
 Leu Thr Leu Ser Pro His Val Glu Leu Ser Gly Glu Ala Ala Tyr Glu  
 900 905 910  
 Leu Arg Gly Ser Ala His Ile Tyr Asn Val Asp Cys Gly Leu Arg Tyr  
 915 920 925  
 Ser Phe  
 930

## (2) INFORMATION FOR SEQ ID NO:17:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 840 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

GAAGACAATA	TAAGGTACCG	TCATAACAGC	GGGGGTTATG	CACTAGGGGAT	CACAGCAACA	60
ACTCCTGCCG	AGGATCAGCT	TACTTTTGCC	TTCTGCCAGC	TCTTTGCTAG	AGATCGCAAT	120
CATATTACAG	GTAAGAACCA	CGGAGATACT	TACGGTGCCT	CTTTGTATTT	CCACCATACA	180
GAAGGGCTCT	TCGACATCGC	CAATTTCTCT	TGGGGAAAAG	CAACCCGAGC	TCCCTGGGTG	240
CTCTCTGAGA	TCTCCCAGAT	CATTCCTTTA	TCGTTTCGATG	CTAAATTCAG	TTATCTCCAT	300
ACAGACAACC	ACATGAAGAC	ATATTATACC	GATAACTCTA	TCATCAAGGG	TTCTTGGAGA	360
AACGATGCCT	TCTGTGCAGA	TCTTGGAGCT	AGCCTGCCTT	TTGTTATTTC	CGTTCCGTAT	420
CTTCTGAAAG	AAGTCGAACC	TTTTGTCAAA	GTACAGTATA	TCTATGCGCA	TCAGCAAGAC	480
TTCTACGAGC	GTCATGCTGA	AGGACGCGCT	TTCAATAAAA	GCGAGCTTAT	CAACGTAGAG	540
ATTCCTATAG	GCGTCACCTT	CGAAAGAGAC	TCAAATCAG	AAAAGGGAAC	TTACGATCTT	600
ACTCTTATGT	ATATACTCGA	TGCTTACCGA	CGCAATCCTA	AATGTCAAAC	TTCCCTAATA	660
GCTAGCGATG	CTAACTGGAT	GGCCTATGGT	ACCAACCTCG	CACGACAAGG	TTTTTCTGTT	720
CGTGCTGCGA	ACCATTTCCA	AGTGAACCCC	CACATGGAAA	TCTTCGGTCA	ATTCGCTTTT	780
GAAGTACGAA	GTTCTTCACG	AAATTATAAT	ACAAACCTAG	GCTCTAAGTT	TTGTTTCTAG	840

## (2) INFORMATION FOR SEQ ID NO:18:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 279 amino acids  
 (B) TYPE: amino acid

(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

```

Glu Asp Asn Ile Arg Tyr Arg His Asn Ser Gly Gly Tyr Ala Leu Gly
 1           5           10           15
Ile Thr Ala Thr Thr Pro Ala Glu Asp Gln Leu Thr Phe Ala Phe Cys
      20           25           30
Gln Leu Phe Ala Arg Asp Arg Asn His Ile Thr Gly Lys Asn His Gly
      35           40           45
Asp Thr Tyr Gly Ala Ser Leu Tyr Phe His His Thr Glu Gly Leu Phe
      50           55           60
Asp Ile Ala Asn Phe Leu Trp Gly Lys Ala Thr Arg Ala Pro Trp Val
      65           70           75           80
Leu Ser Glu Ile Ser Gln Ile Ile Pro Leu Ser Phe Asp Ala Lys Phe
      85           90           95
Ser Tyr Leu His Thr Asp Asn His Met Lys Thr Tyr Tyr Thr Asp Asn
      100          105          110
Ser Ile Ile Lys Gly Ser Trp Arg Asn Asp Ala Phe Cys Ala Asp Leu
      115          120          125
Gly Ala Ser Leu Pro Phe Val Ile Ser Val Pro Tyr Leu Leu Lys Glu
      130          135          140
Val Glu Pro Phe Val Lys Val Gln Tyr Ile Tyr Ala His Gln Gln Asp
      145          150          155          160
Phe Tyr Glu Arg His Ala Glu Gly Arg Ala Phe Asn Lys Ser Glu Leu
      165          170          175
Ile Asn Val Glu Ile Pro Ile Gly Val Thr Phe Glu Arg Asp Ser Lys
      180          185          190
Ser Glu Lys Gly Thr Tyr Asp Leu Thr Leu Met Tyr Ile Leu Asp Ala
      195          200          205
Tyr Arg Arg Asn Pro Lys Cys Gln Thr Ser Leu Ile Ala Ser Asp Ala
      210          215          220
Asn Trp Met Ala Tyr Gly Thr Asn Leu Ala Arg Gln Gly Phe Ser Val
      225          230          235          240
Arg Ala Ala Asn His Phe Gln Val Asn Pro His Met Glu Ile Phe Gly
      245          250          255
Gln Phe Ala Phe Glu Val Arg Ser Ser Arg Asn Tyr Asn Thr Asn
      260          265          270
Leu Gly Ser Lys Phe Cys Phe
      275

```

(2) INFORMATION FOR SEQ ID NO:19:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1545 base pairs  
(B) TYPE: nucleic acid  
(C) STRANDEDNESS: single  
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: Genomic DNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:

ATGACCATAC TTCGAAATTT TCTTACCTGC TCGGCTTTAT TCCTCGCTCT CCCTGCAGCA

GCACAAGTTG	TATATCTTCA	TGAAAGTGAT	GGTTATAACG	GTGCTATCAA	TAATAAAAGC	120
TTAGAACCTA	AAATTACCTG	TTATCCAGAA	GGAACCTCTT	ACATCTTTCT	AGATGACGTG	180
AGGATTTCCA	ACGTTAAGCA	TGATCAAGAA	GATGCTGGGG	TTTTTATAAA	TCGATCTGGG	240
AATCTTTTTT	TCATGGGCAA	CCGTTGCAAC	TTCACTTTTT	ACAACCTTAT	GACCGAGGGT	300
TTTGGCGCTG	CCATTTGAA	CCGCGTTGGA	GACACCACTC	TCACTCTCTC	TAATTTTTCT	360
TACTTAACGT	TCACCTCAGC	ACCTCTACTA	CCTCAAGGAC	AAGGAGCGAT	TTATAGTCTT	420
GGTTCCGTGA	TGATCGAAAA	TAGTGAGGAA	GTGACTTTCT	GTGGGAACTA	CTCTTCGTGG	480
AGTGGAGCTG	CGATTTATAC	TCCCTACCTT	TTAGGTTCTA	AGGCGAGTCG	TCCTTCAGTA	540
AATCTCAGCG	GGAACCGCTA	CCTGGTGTTC	AGAGACTATG	TGAGCCAAGG	TTATGGCGGC	600
GCCGTATCTA	CCCACAATCT	CACACTCACG	ACTCGAGGAC	CTTCGTGTTT	TGAAAATAAT	660
CATGCTTATC	ATGACGTGAA	TAGTAATGGA	GGAGCCATTG	CCATTGCTCC	TGGAGGATCG	720
ATCTCTATAT	CCGTGAAAAG	CGGAGATCTC	ATCTTCAAAG	GAAATACAGC	ATCACAAGAC	780
GGAAATACAA	TACACAATC	CATCCATCTG	CAATCTGGAG	CACAGTTTAA	GAACCTACGT	840
GCTGTTTCAG	AATCCGGAGT	TTATTTCTAT	GATCCTATAA	GCCATAGCGA	GTCGCATAAA	900
ATTACAGATC	TTGTAATCAA	TGCTCCTGAA	GGAAAGGAAA	CTTATGAAGG	AACAATTAGC	960
TTCTCAGGAC	TATGCCTGGA	TGATCATGAA	GTTTGTGCGG	AAAATCTTAC	TTCCACAATC	1020
CTACAAGATG	TCACATTAGC	AGGAGGAACT	CTCTCTCTAT	CGGATGGGGT	TACCTTGCAA	1080
CTGCATTCTT	TTAAGCAGGA	AGCAAGCTCT	ACGCTTACTA	TGTCTCCAGG	AACCACTCTG	1140
CTCTGCTCAG	GAGATGCTCG	GGTTCAGAAT	CTGCACATCC	TGATTGAAGA	TAQCGACAAC	1200
TTTGTTTCTG	TAAGGATTCTG	CGCCGAGGAC	AAGGATGCTC	TTGTCTCATT	AGAAAACTT	1260
AAAGTTGCCT	TTGAGGCTTA	TTGGTCCGTC	TATGACTTTC	CTCAATTTAA	GGAAGCCTTT	1320
ACGATTCTCT	TTCTTGAAT	TCTAGGGCCT	TCTTTTGACA	GTCTTCTCCT	AGGGGAGACC	1380
ACTTTGGAGA	GAACCCAAGT	CACAACAGAG	AATGACGCCG	TTGAGGTTT	CTGGTCCCTA	1440
AGCTGGGAAG	AGTACCCCCC	TTCTCTGGAT	AAAGACAGAA	GGATCACACC	AACTAAGAAA	1500
ACTGTTTTCC	TCACTTGGA	TCCTGAGATC	ACTTCTACGC	CATAA		1545

## (2) INFORMATION FOR SEQ ID NO:20:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 514 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:

Met	Thr	Ile	Leu	Arg	Asn	Phe	Leu	Thr	Cys	Ser	Ala	Leu	Phe	Leu	Ala
1				5					10					15	
Leu	Pro	Ala	Ala	Ala	Gln	Val	Val	Tyr	Leu	His	Glu	Ser	Asp	Gly	Tyr
			20					25					30		
Asn	Gly	Ala	Ile	Asn	Asn	Lys	Ser	Leu	Glu	Pro	Lys	Ile	Thr	Cys	Tyr
			35				40						45		
Pro	Glu	Gly	Thr	Ser	Tyr	Ile	Phe	Leu	Asp	Asp	Val	Arg	Ile	Ser	Asn
			50				55					60			
Val	Lys	His	Asp	Gln	Glu	Asp	Ala	Gly	Val	Phe	Ile	Asn	Arg	Ser	Gly
65				70					75				80		
Asn	Leu	Phe	Phe	Met	Gly	Asn	Arg	Cys	Asn	Phe	Thr	Phe	His	Asn	Leu
			85					90					95		
Met	Thr	Glu	Gly	Phe	Gly	Ala	Ala	Ile	Ser	Asn	Arg	Val	Gly	Asp	Thr
			100					105					110		
Thr	Leu	Thr	Leu	Ser	Asn	Phe	Ser	Tyr	Leu	Thr	Phe	Thr	Ser	Ala	Pro
			115				120						125		
Leu	Leu	Pro	Gln	Gly	Gln	Gly	Ala	Ile	Tyr	Ser	Leu	Gly	Ser	Val	Met
			130				135					140			
Ile	Glu	Asn	Ser	Glu	Glu	Val	Thr	Phe	Cys	Gly	Asn	Tyr	Ser	Ser	Trp



145                      150                      155                      160  
 Ser Gly Ala Ala Ile Tyr Thr Pro Tyr Leu Leu Gly Ser Lys Ala Ser  
                                  165                      170                      175  
 Arg Pro Ser Val Asn Leu Ser Gly Asn Arg Tyr Leu Val Phe Arg Asp  
                                  180                      185                      190  
 Tyr Val Ser Gln Gly Tyr Gly Gly Ala Val Ser Thr His Asn Leu Thr  
                                  195                      200                      205  
 Leu Thr Thr Arg Gly Pro Ser Cys Phe Glu Asn Asn His Ala Tyr His  
                                  210                      215                      220  
 Asp Val Asn Ser Asn Gly Gly Ala Ile Ala Ile Ala Pro Gly Gly Ser  
 225                                   230                      235                      240  
 Ile Ser Ile Ser Val Lys Ser Gly Asp Leu Ile Phe Lys Gly Asn Thr  
                                  245                      250                      255  
 Ala Ser Gln Asp Gly Asn Thr Ile His Asn Ser Ile His Leu Gln Ser  
                                  260                      265                      270  
 Gly Ala Gln Phe Lys Asn Leu Arg Ala Val Ser Glu Ser Gly Val Tyr  
                                  275                      280                      285  
 Phe Tyr Asp Pro Ile Ser His Ser Glu Ser His Lys Ile Thr Asp Leu  
                                  290                      295                      300  
 Val Ile Asn Ala Pro Glu Gly Lys Glu Thr Tyr Glu Gly Thr Ile Ser  
 305                                   310                      315                      320  
 Phe Ser Gly Leu Cys Leu Asp Asp His Glu Val Cys Ala Glu Asn Leu  
                                  325                      330                      335  
 Thr Ser Thr Ile Leu Gln Asp Val Thr Leu Ala Gly Gly Thr Leu Ser  
                                  340                      345                      350  
 Leu Ser Asp Gly Val Thr Leu Gln Leu His Ser Phe Lys Gln Glu Ala  
                                  355                      360                      365  
 Ser Ser Thr Leu Thr Met Ser Pro Gly Thr Thr Leu Leu Cys Ser Gly  
                                  370                      375                      380  
 Asp Ala Arg Val Gln Asn Leu His Ile Leu Ile Glu Asp Thr Asp Asn  
 385                                   390                      395                      400  
 Phe Val Pro Val Arg Ile Arg Ala Glu Asp Lys Asp Ala Leu Val Ser  
                                  405                      410                      415  
 Leu Glu Lys Leu Lys Val Ala Phe Glu Ala Tyr Trp Ser Val Tyr Asp  
                                  420                      425                      430  
 Phe Pro Gln Phe Lys Glu Ala Phe Thr Ile Pro Leu Leu Glu Leu Leu  
                                  435                      440                      445  
 Gly Pro Ser Phe Asp Ser Leu Leu Leu Gly Glu Thr Thr Leu Glu Arg  
                                  450                      455                      460  
 Thr Gln Val Thr Thr Glu Asn Asp Ala Val Arg Gly Phe Trp Ser Leu  
 465                                   470                      475                      480  
 Ser Trp Glu Glu Tyr Pro Pro Ser Leu Asp Lys Asp Arg Arg Ile Thr  
                                  485                      490                      495  
 Pro Thr Lys Lys Thr Val Phe Leu Thr Trp Asn Pro Glu Ile Thr Ser  
                                  500                      505                      510  
 Thr Pro

## (2) INFORMATION FOR SEQ ID NO:21:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 787 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:21:

ATGAAAACGT	CTATTCGTAA	GTTCTTAATT	TCTACCACAC	TGGCGCCATG	TTTTGCTTCA	60
ACAGCGTTTA	CTGTAGAAGT	TATCATGCCT	TCCGAGAACT	TTGATGGATC	GAGTGGGAAG	120
ATTTTTCCTT	ACACAACACT	TTCTGATCCT	AGAGGGACAC	TCTGTATTTT	TTCAGGGGAT	180
CTCTACATTG	CGAATCTTGA	TAATGCCATA	TCCAGAACCT	CTTCCAGTTG	CTTTAGCAAT	240
AGGGCGGGAG	CACTACAAAT	CTTAGGAAAA	GGTGGGGTTT	TCTCCTTCTT	AAATATCCGT	300
TCTTCAGCTG	ACGGAGCCGC	GATTAGTAGT	GTAATCACCC	AAAATCCTGA	ACTATGTCCC	360
TTGAGTTTTT	CAGGATTTAG	TCAGATGATC	TTCGATAACT	GTGAATCTTT	GACTTCAGAT	420
ACCTCAGCGA	GTAATGTCAT	ACCTCACGCA	TCGGCGATTT	ACGCTACAAC	GCCCATGCTC	480
TTTACAAACA	ATGACTCCAT	ACTATTCCTA	TACAACCGTT	CTGCAGGATT	TGGAGCTGCC	540
ATTCGAGGCA	CAAGCATCAC	AATAGAAAAT	ACGAAAAAGA	GCCTTCTCTT	TAATGGTAAT	600
GGATCCATCT	CTAATGGAGG	GGCCCTCACG	GGATCTGCAG	CGATCAACCT	CATCAACAAT	660
AGCGCTCCTG	TGATTTTCTC	AACGAATGCT	ACAGGGATCT	ATGGTGGGGC	TATTTACCTT	720
ACCGGAGGAT	CTATGCTCAC	CTCTGGGAAC	CTCTCAGGAG	TCTTGTTTCGT	TTATAATAGC	780
TCGCGCT						787

## (2) INFORMATION FOR SEQ ID NO:22:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 262 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

Met	Lys	Thr	Ser	Ile	Arg	Lys	Phe	Leu	Ile	Ser	Thr	Thr	Leu	Ala	Pro
1				5				10					15		
Cys	Phe	Ala	Ser	Thr	Ala	Phe	Thr	Val	Glu	Val	Ile	Met	Pro	Ser	Glu
			20					25					30		
Asn	Phe	Asp	Gly	Ser	Ser	Gly	Lys	Ile	Phe	Pro	Tyr	Thr	Thr	Leu	Ser
			35				40					45			
Asp	Pro	Arg	Gly	Thr	Leu	Cys	Ile	Phe	Ser	Gly	Asp	Leu	Tyr	Ile	Ala
			50			55				60					
Asn	Leu	Asp	Asn	Ala	Ile	Ser	Arg	Thr	Ser	Ser	Ser	Cys	Phe	Ser	Asn
65					70				75					80	
Arg	Ala	Gly	Ala	Leu	Gln	Ile	Leu	Gly	Lys	Gly	Gly	Val	Phe	Ser	Phe
				85				90					95		
Leu	Asn	Ile	Arg	Ser	Ser	Ala	Asp	Gly	Ala	Ala	Ile	Ser	Ser	Val	Ile
			100					105				110			
Thr	Gln	Asn	Pro	Glu	Leu	Cys	Pro	Leu	Ser	Phe	Ser	Gly	Phe	Ser	Gln
			115				120					125			
Met	Ile	Phe	Asp	Asn	Cys	Glu	Ser	Leu	Thr	Ser	Asp	Thr	Ser	Ala	Ser
			130			135				140					
Asn	Val	Ile	Pro	His	Ala	Ser	Ala	Ile	Tyr	Ala	Thr	Thr	Pro	Met	Leu
145					150				155					160	
Phe	Thr	Asn	Asn	Asp	Ser	Ile	Leu	Phe	Gln	Tyr	Asn	Arg	Ser	Ala	Gly
				165				170					175		
Phe	Gly	Ala	Ala	Ile	Arg	Gly	Thr	Ser	Ile	Thr	Ile	Glu	Asn	Thr	Lys
			180				185					190			
Lys	Ser	Leu	Leu	Phe	Asn	Gly	Asn	Gly	Ser	Ile	Ser	Asn	Gly	Gly	Ala
			195			200					205				
Leu	Thr	Gly	Ser	Ala	Ala	Ile	Asn	Leu	Ile	Asn	Asn	Ser	Ala	Pro	Val
			210			215				220					

Ile Phe Ser Thr Asn Ala Thr Gly Ile Tyr Gly Gly Ala Ile Tyr Leu  
 225 230 235 240  
 Thr Gly Gly Ser Met Leu Thr Ser Gly Asn Leu Ser Gly Val Leu Phe  
 245 250 255  
 Val Tyr Asn Ser Ser Arg  
 260

## (2) INFORMATION FOR SEQ ID NO:23:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2838 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: Genomic DNA

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:

ATGAAGACTT CAGTTTCTAT GTTGTGGGCC CTGCTTTGCT CGGGGGCTAG CTCIATTGTA 60  
 CTCCATGCCG CAACCACTCC ACTAAATCCT GAAGATGGGT TTATTGGGGA GGGCAATACA 120  
 AATACTTTT CTCCGAAATC TACAACGGAT GCTGCAGGAA CTACCTACTC TCTCACAGGA 180  
 GAGGTTCTGT TTATAGATCC GGGGAAAGGT GGTTC AATTA CAGGAAC TTG CTTTGTAGAA 240  
 ACTGCTGGCG ATCTTACATT TTTAGGTAAT GGAAATACCC TAAAGTTCCT GTCGGTAGAT 300  
 GCAGGTGCTA ATATCGCGGT TGCTCATGTA CAAGGAAGTA AGAATTTAAG CTCACAGAT 360  
 TTCCPTTCTC TGGTGATCAC AGAATCTCCA AAATCCGCTG TTAGTACAGG AAAAGGTAGC 420  
 CTAGTCAGTT CAGGTGCAGT CCAACTGCAA GATATAAACA CTCTAGTTCT TACAAGCAAT 480  
 GCCTCTGTCT AAGATGGTGG CGTGATTAAA GGAACTCCT GCTTGATTCA GGGAAATCAA 540  
 AATAGTGCGA TTTTGGACA AAATACATCT TCGAAAAAG GAGGGGCGAT CTCCACGACT 600  
 CAAGGACTCA CCATAGAGAA TAACCTTAGGG ACGCTAAGT TCAATGAAAA CAAAGCAGTG 660  
 ACCTCAGGAG GCGCCTTAGA TTTAGGAGCC GCGTCTACAT TCACTGCGAA CCATGAGTTG 720  
 ATATTTTTCAC AAAATAAGAC TTCTGGGAAT GCTGCAAATG GCGGAGCCAT AAATTGCTCA 780  
 GGCGACCTAA CATTTACTGA TAACACTTCT TTGTTACTTC AAGAAAATAG CACAATGCAG 840  
 GATGGTGGAG CTTTGTGTAG CACAGGAACC ATAAGCATTA CCGGTAGTGA TTCTATCAAT 900  
 GTGATAGGAA ATACTTCAGG ACAAAGGGA GGAGCGATTT CTGCAGCTTC TCTCAAGATT 960  
 TTGGGAGGGC AGGGAGGCGC TCTCTTTTCT AATAACGTAG TGACTCATGC CACCCCTCTA 1020  
 GGAGGTGCCA TTTTATCAA CACAGGAGGA TCCTTGCGAGC TCTTCACTCA AGGAGGGGAT 1080  
 ATCGTATTCG AGGGGAATCA GGTCACATA ACAGCTCCAA ATGCTACCAC TAAGAGAAAT 1140  
 GTAATTCACC TCGAGAGCAC CGCGAAGTGG ACGGGACTTG CTGCAAGTCA AGGTAACGCT 1200  
 ATCTATTTCT ATGATCCCAT TACCACCAAC GATACGGGAG CAAGCGATAA CTTACGTATC 1260  
 AATGAGGTCA GTGCAATCA AAAGCTCTCG GGATCTATAG TATTTTCTGG AGAGAGATTG 1320  
 TCGACAGCAG AAGCTATAGC TGAAAATCTT ACTTCGAGGA TCAACCAGCC TGTCATTTA 1380  
 GTAGAGGGGA GCTTAGAACT TAAACAGGGA GTGACCTTGA TCACACAAGG ATTCTCGCAG 1440  
 GAGCCAGAAT CCACGCTTCT TTTGGATTG GGGACCTCAT TACAAGCTTC TACAGAAGAT 1500  
 ATCGTCATCA CAAATTCATC TATAAATGCC GATACCATTT ACGGAAAGAA TCCAATCAAT 1560  
 ATTGTAGCTT CAGCAGCGAA TAAGAACATT ACCCTAACAG GAACCTTAGC ACTTGTAAAT 1620  
 GCAGATGGAG CTTTGTATGA GAACCATACC TTGCAAGACT CTCAAGATTA TAGCTTTGTA 1680  
 AAGTTATCTC CAGGAGCGGG AGGGACTATA ATTACTCAAG ATGCTTCTCA GAAGCTTCTT 1740  
 GAAGTAGCTC CTTCTAGACC ACATTATGGC TATCAAGGAC ATTGGAATGT GCAAGTCATC 1800  
 CCAGGAACGG GAACTCAACC GAGCCAGGCA AATTTAGAAAT GGGTGCGGAC AGGATACCTT 1860  
 CCGAATCCC GAAAGCAAGG ATTTTGTAGT CCAATAGACC TGTGGGGTTC TTTTGTGTAT 1920  
 CAGCGTGGCTA TCCAAGAAAT CATGGTAAAT AGTAGCCAAA TCTTATGTCA GGAACGGGGA 1980  
 GTCTGGGGAG CTGGAATTGC TAATTTCTTA CATAGAGATA AAATTAATGA GCACGGCTAT 2040  
 CGCCATAGCG GTGTGGTTA TCTTGTGGGA GTTGGCACTC ATGCTTTTTC TGATGCTACG 2100  
 ATAAATGCGG CTTTTTGCCA GCTCTTCAGT AGAGATAAAG ACTACGTAGT ATCCAAAAAT 2160  
 CATGGAATA GCTACTCAGG GGTCTGATTT CTTGAGGATA CCCTAGAGTT TAGAAGTCCA 2220  
 CAGGGATTCT ATACTGATAG CTCCTCAGAA GCTTGCTGTA ACCAAGTCGT CACTATAGAT 2280

ATGCAGTTGT	CTTACAGCCA	TAGAAATAAT	GATATGAAAA	CCAAATACAC	GACATATCCA	2340
GAAGCTCAGG	GATCTTGGGC	AAATGATGTT	TTTGGTCTTG	AGTTTGGAGC	GACTACATAC	2400
TACTACCCTA	ACAGTACTTT	TTTATTTGAT	TACTACTCTC	CGTTTCTCAG	GCTGCAGTGC	2460
ACCTATGCTC	ACCAGGAAGA	CTTCAAAGAG	ACAGGAGGTG	AGGTTTCGTCA	CTTTACTAGC	2520
GGAGATCTTT	TCAATTTAGC	AGTTCCTATT	GGCGTGAAGT	TTGAGAGATT	TTCAGACTGT	2580
AAAAGGGGAT	CTTATGAACT	TACCCTTGCT	TATGTTTCCTG	ATGTGATTCTG	CAAAGATCCC	2640
AAGAGCACGG	CAACATTGGC	TAGTGGAGCT	ACGTGGAGCA	CCCACGGAAA	CAATCTCTCC	2700
AGACAAGGAT	TACAACGCG	TTTAGGGAAC	CACGTGTCTCA	TAAATCCTGG	AATTGAGGTG	2760
TTCAGTCACG	GAGCTATTGA	ATTGCGGGGA	TCCTCTCGTA	ATTATAACAT	CAATCTCGGG	2820
GGTAAATACC	GATTTTAA					2838

## (2) INFORMATION FOR SEQ ID NO:24:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 946 amino acids  
 (B) TYPE: amino acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: peptide

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:24:

Met	Lys	Thr	Ser	Val	Ser	Met	Leu	Leu	Ala	Leu	Leu	Cys	Ser	Gly	Ala	1	5	10	15
Ser	Ser	Ile	Val	Leu	His	Ala	Ala	Thr	Thr	Pro	Leu	Asn	Pro	Glu	Asp	20	25	30	
Gly	Phe	Ile	Gly	Glu	Gly	Asn	Thr	Asn	Thr	Phe	Ser	Pro	Lys	Ser	Thr	35	40	45	
Thr	Asp	Ala	Ala	Gly	Thr	Thr	Tyr	Ser	Leu	Thr	Gly	Glu	Val	Leu	Phe	50	55	60	
Ile	Asp	Pro	Gly	Lys	Gly	Gly	Ser	Ile	Thr	Gly	Thr	Cys	Phe	Val	Glu	65	70	75	80
Thr	Ala	Gly	Asp	Leu	Thr	Phe	Leu	Gly	Asn	Gly	Asn	Thr	Leu	Lys	Phe	85	90	95	
Leu	Ser	Val	Asp	Ala	Gly	Ala	Asn	Ile	Ala	Val	Ala	His	Val	Gln	Gly	100	105	110	
Ser	Lys	Asn	Leu	Ser	Phe	Thr	Asp	Phe	Leu	Ser	Leu	Val	Ile	Thr	Glu	115	120	125	
Ser	Pro	Lys	Ser	Ala	Val	Ser	Thr	Gly	Lys	Gly	Ser	Leu	Val	Ser	Ser	130	135	140	
Gly	Ala	Val	Gln	Leu	Gln	Asp	Ile	Asn	Thr	Leu	Val	Leu	Thr	Ser	Asn	145	150	155	160
Ala	Ser	Val	Glu	Asp	Gly	Gly	Val	Ile	Lys	Gly	Asn	Ser	Cys	Leu	Ile	165	170	175	
Gln	Gly	Ile	Lys	Asn	Ser	Ala	Ile	Phe	Gly	Gln	Asn	Thr	Ser	Ser	Lys	180	185	190	
Lys	Gly	Gly	Ala	Ile	Ser	Thr	Thr	Gln	Gly	Leu	Thr	Ile	Glu	Asn	Asn	195	200	205	
Leu	Gly	Thr	Leu	Lys	Phe	Asn	Glu	Asn	Lys	Ala	Val	Thr	Ser	Gly	Gly	210	215	220	
Ala	Leu	Asp	Leu	Gly	Ala	Ala	Ser	Thr	Phe	Thr	Ala	Asn	His	Glu	Leu	225	230	235	240
Ile	Phe	Ser	Gln	Asn	Lys	Thr	Ser	Gly	Asn	Ala	Ala	Asn	Gly	Gly	Ala	245	250	255	
Ile	Asn	Cys	Ser	Gly	Asp	Leu	Thr	Phe	Thr	Asp	Asn	Thr	Ser	Leu	Leu	260	265	270	

Leu Gln Glu Asn Ser Thr Met Gln Asp Gly Gly Ala Leu Cys Ser Thr  
 275 280 285  
 Gly Thr Ile Ser Ile Thr Gly Ser Asp Ser Ile Asn Val Ile Gly Asn  
 290 295 300  
 Thr Ser Gly Gln Lys Gly Gly Ala Ile Ser Ala Ala Ser Leu Lys Ile  
 305 310 315 320  
 Leu Gly Gly Gln Gly Gly Ala Leu Phe Ser Asn Asn Val Val Thr His  
 325 330 335  
 Ala Thr Pro Leu Gly Gly Ala Ile Phe Ile Asn Thr Gly Gly Ser Leu  
 340 345 350  
 Gln Leu Phe Thr Gln Gly Gly Asp Ile Val Phe Glu Gly Asn Gln Val  
 355 360 365  
 Thr Thr Thr Ala Pro Asn Ala Thr Thr Lys Arg Asn Val Ile His Leu  
 370 375 380  
 Glu Ser Thr Ala Lys Trp Thr Gly Leu Ala Ala Ser Gln Gly Asn Ala  
 385 390 395 400  
 Ile Tyr Phe Tyr Asp Pro Ile Thr Thr Asn Asp Thr Gly Ala Ser Asp  
 405 410 415  
 Asn Leu Arg Ile Asn Glu Val Ser Ala Asn Gln Lys Leu Ser Gly Ser  
 420 425 430  
 Ile Val Phe Ser Gly Glu Arg Leu Ser Thr Ala Glu Ala Ile Ala Glu  
 435 440 445  
 Asn Leu Thr Ser Arg Ile Asn Gln Pro Val Thr Leu Val Glu Gly Ser  
 450 455 460  
 Leu Glu Leu Lys Gln Gly Val Thr Leu Ile Thr Gln Gly Phe Ser Gln  
 465 470 475 480  
 Glu Pro Glu Ser Thr Leu Leu Leu Asp Leu Gly Thr Ser Leu Gln Ala  
 485 490 495  
 Ser Thr Glu Asp Ile Val Ile Thr Asn Ser Ser Ile Asn Ala Asp Thr  
 500 505 510  
 Ile Tyr Gly Lys Asn Pro Ile Asn Ile Val Ala Ser Ala Asn Lys  
 515 520 525  
 Asn Ile Thr Leu Thr Gly Thr Leu Ala Leu Val Asn Ala Asp Gly Ala  
 530 535 540  
 Leu Tyr Glu Asn His Thr Leu Gln Asp Ser Gln Asp Tyr Ser Phe Val  
 545 550 555 560  
 Lys Leu Ser Pro Gly Ala Gly Gly Thr Ile Ile Thr Gln Asp Ala Ser  
 565 570 575  
 Gln Lys Leu Leu Glu Val Ala Pro Ser Arg Pro His Tyr Gly Tyr Gln  
 580 585 590  
 Gly His Trp Asn Val Gln Val Ile Pro Gly Thr Gly Thr Gln Pro Ser  
 595 600 605  
 Gln Ala Asn Leu Glu Trp Val Arg Thr Gly Tyr Leu Pro Asn Pro Glu  
 610 615 620  
 Arg Gln Gly Phe Leu Val Pro Asn Ser Leu Trp Gly Ser Phe Val Asp  
 625 630 635 640  
 Gln Arg Ala Ile Gln Glu Ile Met Val Asn Ser Ser Gln Ile Leu Cys  
 645 650 655  
 Gln Glu Arg Gly Val Trp Gly Ala Gly Ile Ala Asn Phe Leu His Arg  
 660 665 670  
 Asp Lys Ile Asn Glu His Gly Tyr Arg His Ser Gly Val Gly Tyr Leu  
 675 680 685  
 Val Gly Val Gly Thr His Ala Phe Ser Asp Ala Thr Ile Asn Ala Ala  
 690 695 700  
 Phe Cys Gln Leu Phe Ser Arg Asp Lys Asp Tyr Val Val Ser Lys Asn  
 705 710 715 720  
 His Gly Thr Ser Tyr Ser Gly Val Val Phe Leu Glu Asp Thr Leu Glu

725 730 735  
 Phe Arg Ser Pro Gln Gly Phe Tyr Thr Asp Ser Ser Ser Glu Ala Cys  
 740 745 750  
 Cys Asn Gln Val Val Thr Ile Asp Met Gln Leu Ser Tyr Ser His Arg  
 755 760 765  
 Asn Asn Asp Met Lys Thr Lys Tyr Thr Thr Tyr Pro Glu Ala Gln Gly  
 770 775 780  
 Ser Trp Ala Asn Asp Val Phe Gly Leu Glu Phe Gly Ala Thr Thr Tyr  
 785 790 795 800  
 Tyr Tyr Pro Asn Ser Thr Phe Leu Phe Asp Tyr Tyr Ser Pro Phe Leu  
 805 810 815  
 Arg Leu Gln Cys Thr Tyr Ala His Gln Glu Asp Phe Lys Glu Thr Gly  
 820 825 830  
 Gly Glu Val Arg His Phe Thr Ser Gly Asp Leu Phe Asn Leu Ala Val  
 835 840 845  
 Pro Ile Gly Val Lys Phe Glu Arg Phe Ser Asp Cys Lys Arg Gly Ser  
 850 855 860  
 Tyr Glu Leu Thr Leu Ala Tyr Val Pro Asp Val Ile Arg Lys Asp Pro  
 865 870 875 880  
 Lys Ser Thr Ala Thr Leu Ala Ser Gly Ala Thr Trp Ser Thr His Gly  
 885 890 895  
 Asn Asn Leu Ser Arg Gln Gly Leu Gln Leu Arg Leu Gly Asn His Cys  
 900 905 910  
 Leu Ile Asn Pro Gly Ile Glu Val Phe Ser His Gly Ala Ile Glu Leu  
 915 920 925  
 Arg Gly Ser Ser Arg Asn Tyr Asn Ile Asn Leu Gly Gly Lys Tyr Arg  
 930 935 940  
 Phe  
 945

## (2) INFORMATION FOR SEQ ID NO:25:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 3000 base pairs  
 (B) TYPE: nucleic acid  
 (C) STRANDEDNESS: single  
 (D) TOPOLOGY: linear

## (ix) FEATURE:

- (A) NAME/KEY: Coding Sequence  
 (B) LOCATION: 259...3000  
 (D) OTHER INFORMATION:

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:

ATCAGGTGAT AAAAGTTCCT CGTTAGCTAG TGACTGTAGG TGACATGAGA AAGCTAACAC 60  
 GGAGGAAACT AAAACCCAAG GAATCGAAGT CTTCATGGTA ATGCTTTTGT TTTTATAGAGA 120  
 ACTATTCGCA TCAATATAGA AACAAAATAA GTAAATCAAG TTAAAGATGA CAAAACAGCT 180  
 GTCAAGAATT TTTATCTTGA CTCTCTGAGT TTTCTATTTT ATATGACGCA AGTAAGAATT 240  
 TAATAATAAA GTGGGTTT ATG AAA TCG CAA TTT TCC TGG TTA GTG CTC TCT 291  
 Met Lys Ser Gln Phe Ser Trp Leu Val Leu Ser  
 1 5 10

TCG ACA TTG GCA TGT TTT ACT AGT TGT TCC ACT GTT TTT GCT GCA ACT 339  
 Ser Thr Leu Ala Cys Phe Thr Ser Cys Ser Thr Val Phe Ala Ala Thr  
 15 20 25

GCT GAA AAT ATA GGC CCC TCT GAT AGC TTT GAC GGA AGT ACT AAC ACA 387  
 Ala Glu Asn Ile Gly Pro Ser Asp Ser Phe Asp Gly Ser Thr Asn Thr  
 30 35 40

GGC ACC TAT ACT CCT AAA AAT ACG ACT ACT GGA ATA GAC TAT ACT CTG 435  
 Gly Thr Tyr Thr Pro Lys Asn Thr Thr Thr Gly Ile Asp Tyr Thr Leu  
 45 50 55

ACA GGA GAT ATA ACT CTG CAA AAC CTT GGG GAT TCG GCA GCT TTA ACG 483  
 Thr Gly Asp Ile Thr Leu Gln Asn Leu Gly Asp Ser Ala Ala Leu Thr  
 60 65 70 75

AAG GGT TGT TTT TCT GAC ACT ACG GAA TCT TTA AGC TTT GCC GGT AAG 531  
 Lys Gly Cys Phe Ser Asp Thr Thr Glu Ser Leu Ser Phe Ala Gly Lys  
 80 85 90

GGG TAC TCA CTT TCT TTT TTA AAT ATT AAG TCT AGT GCT GAA GGC GCA 579  
 Gly Tyr Ser Leu Ser Phe Leu Asn Ile Lys Ser Ser Ala Glu Gly Ala  
 95 100 105

GCA CTT TCT GTT ACA ACT GAT AAA AAT CTG TCG CTA ACA GGA TTT TCG 627  
 Ala Leu Ser Val Thr Thr Asp Lys Asn Leu Ser Leu Thr Gly Phe Ser  
 110 115 120

AGT CTT ACT TTC TTA GCG GCC CCA TCA TCG GTA ATC ACA ACC CCC TCA 675  
 Ser Leu Thr Phe Leu Ala Ala Pro Ser Ser Val Ile Thr Thr Pro Ser  
 125 130 135

GGA AAA GGT GCA GTT AAA TGT GGA GGG GAT CTT ACA TTT GAT AAC AAT 723  
 Gly Lys Gly Ala Val Lys Cys Gly Gly Asp Leu Thr Phe Asp Asn Asn  
 140 145 150 155

GGA ACT ATT TTA TTT AAA CAA GAT TAC TGT GAG GAA AAT GGC GGA GCC 771  
 Gly Thr Ile Leu Phe Lys Gln Asp Tyr Cys Glu Glu Asn Gly Gly Ala  
 160 165 170

ATT TCT ACC AAG AAT CTT TCT TTG AAA AAC AGC ACG GGA TCG ATT TCT 819  
 Ile Ser Thr Lys Asn Leu Ser Leu Lys Asn Ser Thr Gly Ser Ile Ser  
 175 180 185

TTT GAA GGG AAT AAA TCG AGC GCA ACA GGG AAA AAA GGT GGG GCT ATT 867  
 Phe Glu Gly Asn Lys Ser Ser Ala Thr Gly Lys Lys Gly Gly Ala Ile  
 190 195 200

TGT GCT ACT GGT ACT GTA GAT ATT ACA AAT AAT ACG GCT CCT ACC CTC 915  
 Cys Ala Thr Gly Thr Val Asp Ile Thr Asn Asn Thr Ala Pro Thr Leu  
 205 210 215

TTC TCG AAC AAT ATT GCT GAA GCT GCA GGT GGA GCT ATA AAT AGC ACA 963  
 Phe Ser Asn Asn Ile Ala Glu Ala Ala Gly Gly Ala Ile Asn Ser Thr  
 220 225 230 235

GGA AAC TGT ACA ATT ACA GGG AAT ACG TCT CTT GTA TTT TCT GAA AAT 1011

Gly	Asn	Cys	Thr	Ile	Thr	Gly	Asn	Thr	Ser	Leu	Val	Phe	Ser	Glu	Asn	
				240					245					250		
AGT	GTG	ACA	GCG	ACC	GCA	GGA	AAT	GGA	GGA	GCT	CTT	TCT	GGA	GAT	GCC	1059
Ser	Val	Thr	Ala	Thr	Ala	Gly	Asn	Gly	Gly	Ala	Leu	Ser	Gly	Asp	Ala	
			255				260						265			
GAT	GTT	ACC	ATA	TCT	GGG	AAT	CAG	AGT	GTA	ACT	TTC	TCA	GGA	AAC	CAA	1107
Asp	Val	Thr	Ile	Ser	Gly	Asn	Gln	Ser	Val	Thr	Phe	Ser	Gly	Asn	Gln	
			270				275					280				
GCT	GTA	GCT	AAT	GGC	GGA	GCC	ATT	TAT	GCT	AAG	AAG	CTT	ACA	CTG	GCT	1155
Ala	Val	Ala	Asn	Gly	Gly	Ala	Ile	Tyr	Ala	Lys	Lys	Leu	Thr	Leu	Ala	
	285					290					295					
TCC	GGG	GGG	GGG	GGG	GGT	ATC	TCC	TTT	TCT	AAC	AAT	ATA	GTC	CAA	GGT	1203
Ser	Gly	Gly	Gly	Gly	Gly	Ile	Ser	Phe	Ser	Asn	Asn	Ile	Val	Gln	Gly	
300					305					310				315		
ACC	ACT	GCA	GGT	AAT	GGT	GGA	GCC	ATT	TCT	ATA	CTG	GCA	GCT	GGA	GAG	1251
Thr	Thr	Ala	Gly	Asn	Gly	Gly	Ala	Ile	Ser	Ile	Leu	Ala	Ala	Gly	Glu	
				320					325					330		
TGT	AGT	CTT	TCA	GCA	GAA	GCA	GGG	GAC	ATT	ACC	TTC	AAT	GGG	AAT	GCC	1299
Cys	Ser	Leu	Ser	Ala	Glu	Ala	Gly	Asp	Ile	Thr	Phe	Asn	Gly	Asn	Ala	
			335					340					345			
ATT	GTT	GCA	ACT	ACA	CCA	CAA	ACT	ACA	AAA	AGA	AAT	TCT	ATT	GAC	ATA	1347
Ile	Val	Ala	Thr	Thr	Pro	Gln	Thr	Thr	Lys	Arg	Asn	Ser	Ile	Asp	Ile	
			350				355					360				
GGA	TCT	ACT	GCA	AAG	ATC	ACG	AAT	TTA	CGT	GCA	ATA	TCT	GGG	CAT	AGC	1395
Gly	Ser	Thr	Ala	Lys	Ile	Thr	Asn	Leu	Arg	Ala	Ile	Ser	Gly	His	Ser	
	365					370					375					
ATC	TTT	TTC	TAC	GAT	CCG	ATT	ACT	GCT	AAT	ACG	GCT	GCG	GAT	TCT	ACA	1443
Ile	Phe	Phe	Tyr	Asp	Pro	Ile	Thr	Ala	Asn	Thr	Ala	Ala	Asp	Ser	Thr	
380					385					390					395	
GAT	ACT	TTA	AAT	CTC	AAT	AAG	GCT	GAT	GCA	GGT	AAT	AGT	ACA	GAT	TAT	1491
Asp	Thr	Leu	Asn	Leu	Asn	Lys	Ala	Asp	Ala	Gly	Asn	Ser	Thr	Asp	Tyr	
				400				405					410			
AGT	GGG	TCG	ATT	GTT	TTT	TCT	GGT	GAA	AAG	CTC	TCT	GAA	GAT	GAA	GCA	1539
Ser	Gly	Ser	Ile	Val	Phe	Ser	Gly	Glu	Lys	Leu	Ser	Glu	Asp	Glu	Ala	
			415					420					425			
AAA	GTT	GCA	GAC	AAC	CTC	ACT	TCT	ACG	CTG	AAG	CAG	CCT	GTA	ACT	CTA	1587
Lys	Val	Ala	Asp	Asn	Leu	Thr	Ser	Thr	Leu	Lys	Gln	Pro	Val	Thr	Leu	
			430				435					440				
ACT	GCA	GGA	AAT	TTA	GTA	CTT	AAA	CGT	GGT	GTC	ACT	CTC	GAT	ACG	AAA	1635
Thr	Ala	Gly	Asn	Leu	Val	Leu	Lys	Arg	Gly	Val	Thr	Leu	Asp	Thr	Lys	
	445					450					455					
GGC	TTT	ACT	CAG	ACC	GCG	GGT	TCC	TCT	GTT	ATT	ATG	GAT	GCG	GGC	ACA	1683
Gly	Phe	Thr	Gln	Thr	Ala	Gly	Ser	Ser	Val	Ile	Met	Asp	Ala	Gly	Thr	



460	465	470	475	
ACG TTA AAA GCA AGT ACA GAG GAG GTC ACT TTA ACA GGT CTT TCC ATT				1731
Thr Leu Lys Ala Ser Thr Glu Glu Val Thr Leu Thr Gly Leu Ser Ile				
480		485	490	
CCT GTA GAC TCT TTA GGC GAG GGT AAG AAA GTT GTA ATT GCT GCT TCT				1779
Pro Val Asp Ser Leu Gly Glu Gly Lys Lys Val Val Ile Ala Ala Ser				
495		500	505	
GCA GCA AGT AAA AAT GTA GCC CTT AGT GGT CCG ATT CTT CTT TTG GAT				1827
Ala Ala Ser Lys Asn Val Ala Leu Ser Gly Pro Ile Leu Leu Leu Asp				
510		515	520	
AAC CAA GGG AAT GCT TAT GAA AAT CAC GAC TTA GGA AAA ACT CAA GAC				1875
Asn Gln Gly Asn Ala Tyr Glu Asn His Asp Leu Gly Lys Thr Gln Asp				
525		530	535	
TTT TCA TTT GTG CAG CTC TCT GCT CTG GGT ACT GCA ACA ACT ACA GAT				1923
Phe Ser Phe Val Gln Leu Ser Ala Leu Gly Thr Ala Thr Thr Thr Asp				
540		545	550	555
GTT CCA GCG GTT CCT ACA GTA GCA ACT CCT ACG CAC TAT GGG TAT CAA				1971
Val Pro Ala Val Pro Thr Val Ala Thr Pro Thr His Tyr Gly Tyr Gln				
560		565	570	
GGT ACT TGG GGA ATG ACT TGG GTT GAT GAT ACC GCA AGC ACT CCA AAG				2019
Gly Thr Trp Gly Met Thr Trp Val Asp Asp Thr Ala Ser Thr Pro Lys				
575		580	585	
ACT AAG ACA GCG ACA TTA GCT TGG ACC AAT ACA GGC TAC CTT CCG AAT				2067
Thr Lys Thr Ala Thr Leu Ala Trp Thr Asn Thr Gly Tyr Leu Pro Asn				
590		595	600	
CCT GAG CGT CAA GGA CCT TTA GTT CCT AAT AGC CTT TGG GGA TCT TTT				2115
Pro Glu Arg Gln Gly Pro Leu Val Pro Asn Ser Leu Trp Gly Ser Phe				
605		610	615	
TCA GAC ATC CAA GCG ATT CAA GGT GTC ATA GAG AGA AGT GCT TTG ACT				2163
Ser Asp Ile Gln Ala Ile Gln Gly Val Ile Glu Arg Ser Ala Leu Thr				
620		625	630	635
CTT TGT TCA GAT CGA GGC TTC TGG GCT GCG GGA GTC GCC AAT TTC TTA				2211
Leu Cys Ser Asp Arg Gly Phe Trp Ala Ala Gly Val Ala Asn Phe Leu				
640		645	650	
GAT AAA GAT AAG AAA GGG GAA AAA CGC AAA TAC CGT CAT AAA TCT GGT				2259
Asp Lys Asp Lys Lys Gly Glu Lys Arg Lys Tyr Arg His Lys Ser Gly				
655		660	665	
GGA TAT GCT ATC GGA GGT GCA GCG CAA ACT TGT TCT GAA AAC TTA ATT				2307
Gly Tyr Ala Ile Gly Gly Ala Ala Gln Thr Cys Ser Glu Asn Leu Ile				
670		675	680	
AGC TTT GCC TTT TGC CAA CTC TTT GGT AGC GAT AAA GAT TTC TTA GTC				2355
Ser Phe Ala Phe Cys Gln Leu Phe Gly Ser Asp Lys Asp Phe Leu Val				
685		690	695	

[illegible]

## (2) INFORMATION FOR SEQ ID NO:26:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 914 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: protein

## (v) FRAGMENT TYPE: internal

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:

```

Met Lys Ser Gln Phe Ser Trp Leu Val Leu Ser Ser Thr Leu Ala Cys
 1           5           10           15
Phe Thr Ser Cys Ser Thr Val Phe Ala Thr Ala Glu Asn Ile Gly
      20           25           30
Pro Ser Asp Ser Phe Asp Gly Ser Thr Asn Thr Gly Thr Tyr Thr Pro
    35           40           45
Lys Asn Thr Thr Thr Gly Ile Asp Tyr Thr Leu Thr Gly Asp Ile Thr
   50           55           60
Leu Gln Asn Leu Gly Asp Ser Ala Ala Leu Thr Lys Gly Cys Phe Ser
  65           70           75           80
Asp Thr Thr Glu Ser Leu Ser Phe Ala Gly Lys Gly Tyr Ser Leu Ser
      85           90           95
Phe Leu Asn Ile Lys Ser Ser Ala Glu Gly Ala Ala Leu Ser Val Thr
    100          105          110
Thr Asp Lys Asn Leu Ser Leu Thr Gly Phe Ser Ser Leu Thr Phe Leu
   115          120          125
Ala Ala Pro Ser Ser Val Ile Thr Thr Pro Ser Gly Lys Gly Ala Val
  130          135          140
Lys Cys Gly Gly Asp Leu Thr Phe Asp Asn Asn Gly Thr Ile Leu Phe
 145          150          155          160
Lys Gln Asp Tyr Cys Glu Glu Asn Gly Gly Ala Ile Ser Thr Lys Asn
   165          170          175
Leu Ser Leu Lys Asn Ser Thr Gly Ser Ile Ser Phe Glu Gly Asn Lys
  180          185          190
Ser Ser Ala Thr Gly Lys Lys Gly Gly Ala Ile Cys Ala Thr Gly Thr
   195          200          205
Val Asp Ile Thr Asn Asn Thr Ala Pro Thr Leu Phe Ser Asn Asn Ile
  210          215          220
Ala Glu Ala Ala Gly Gly Ala Ile Asn Ser Thr Gly Asn Cys Thr Ile
 225          230          235          240
Thr Gly Asn Thr Ser Leu Val Phe Ser Glu Asn Ser Val Thr Ala Thr
   245          250          255
Ala Gly Asn Gly Gly Ala Leu Ser Gly Asp Ala Asp Val Thr Ile Ser
  260          265          270
Gln Asn Gln Ser Val Thr Phe Ser Gly Asn Gln Ala Val Ala Asn Gly
   275          280          285
Gly Ala Ile Tyr Ala Lys Lys Leu Thr Leu Ala Ser Gly Gly Gly Gly
  290          295          300
Gly Ile Ser Phe Ser Asn Asn Ile Val Gln Gly Thr Thr Ala Gly Asn
 305          310          315          320
Gly Gly Ala Ile Ser Ile Leu Ala Ala Gly Glu Cys Ser Leu Ser Ala
   325          330          335
Glu Ala Gly Asp Ile Thr Phe Asn Gly Asn Ala Ile Val Ala Thr Thr
 340          345          350

```

Pro Gln Thr Thr Lys Arg Asn Ser Ile Asp Ile Gly Ser Thr Ala Lys  
 355 360 365  
 Ile Thr Asn Leu Arg Ala Ile Ser Gly His Ser Ile Phe Phe Tyr Asp  
 370 375 380  
 Pro Ile Thr Ala Asn Thr Ala Ala Asp Ser Thr Asp Thr Leu Asn Leu  
 385 390 395 400  
 Asn Lys Ala Asp Ala Gly Asn Ser Thr Asp Tyr Ser Gly Ser Ile Val  
 405 410 415  
 Phe Ser Gly Glu Lys Leu Ser Glu Asp Glu Ala Lys Val Ala Asp Asn  
 420 425 430  
 Leu Thr Ser Thr Leu Lys Gln Pro Val Thr Leu Thr Ala Gly Asn Leu  
 435 440 445  
 Val Leu Lys Arg Gly Val Thr Leu Asp Thr Lys Gly Phe Thr Gln Thr  
 450 455 460  
 Ala Gly Ser Ser Val Ile Met Asp Ala Gly Thr Thr Leu Lys Ala Ser  
 465 470 475 480  
 Thr Glu Glu Val Thr Leu Thr Gly Leu Ser Ile Pro Val Asp Ser Leu  
 485 490 495  
 Gly Glu Gly Lys Lys Val Val Ile Ala Ala Ser Ala Ala Ser Lys Asn  
 500 505 510  
 Val Ala Leu Ser Gly Pro Ile Leu Leu Asp Asn Gln Gly Asn Ala  
 515 520 525  
 Tyr Glu Asn His Asp Leu Gly Lys Thr Gln Asp Phe Ser Phe Val Gln  
 530 535 540  
 Leu Ser Ala Leu Gly Thr Ala Thr Thr Thr Asp Val Pro Ala Val Pro  
 545 550 555 560  
 Thr Val Ala Thr Pro Thr His Tyr Gly Tyr Gln Gly Thr Trp Gly Met  
 565 570 575  
 Thr Trp Val Asp Asp Thr Ala Ser Thr Pro Lys Thr Lys Thr Ala Thr  
 580 585 590  
 Leu Ala Trp Thr Asn Thr Gly Tyr Leu Pro Asn Pro Glu Arg Gln Gly  
 595 600 605  
 Pro Leu Val Pro Asn Ser Leu Trp Gly Ser Phe Ser Asp Ile Gln Ala  
 610 615 620  
 Ile Gln Gly Val Ile Glu Arg Ser Ala Leu Thr Leu Cys Ser Asp Arg  
 625 630 635 640  
 Gly Phe Trp Ala Ala Gly Val Ala Asn Phe Leu Asp Lys Asp Lys Lys  
 645 650 655  
 Gly Glu Lys Arg Lys Tyr Arg His Lys Ser Gly Gly Tyr Ala Ile Gly  
 660 665 670  
 Gly Ala Ala Gln Thr Cys Ser Glu Asn Leu Ile Ser Phe Ala Phe Cys  
 675 680 685  
 Gln Leu Phe Gly Ser Asp Lys Asp Phe Leu Val Ala Lys Asn His Thr  
 690 695 700  
 Asp Thr Tyr Ala Gly Ala Phe Tyr Ile Gln His Ile Thr Glu Cys Ser  
 705 710 715 720  
 Gly Phe Ile Gly Cys Leu Leu Asp Lys Leu Pro Gly Ser Trp Ser His  
 725 730 735  
 Lys Pro Leu Val Leu Glu Gly Gln Leu Ala Tyr Ser His Val Ser Asn  
 740 745 750  
 Asp Leu Lys Thr Lys Tyr Thr Ala Tyr Pro Glu Val Lys Gly Ser Trp  
 755 760 765  
 Gly Asn Asn Ala Phe Asn Met Met Leu Gly Ala Ser Ser His Ser Tyr  
 770 775 780  
 Pro Glu Tyr Leu His Cys Phe Asp Thr Tyr Ala Pro Tyr Ile Lys Leu  
 785 790 795 800  
 Asn Leu Thr Tyr Ile Arg Gln Asp Ser Phe Ser Glu Lys Gly Thr Glu

(i) SEQUENCE CHARACTERISTICS:

- (ix) FEATURE:

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:27:

GAT	CCT	AAA	AAT	AAA	GAG	TAC	ACA	GGG	ACC	ATA	CTC	TTT	TCT	GGA	GAA	48
Asp	Pro	Lys	Asn	Lys	Glu	Tyr	Thr	Gly	Thr	Ile	Leu	Phe	Ser	Gly	Glu	
1				5					10					15		
AAG	AGT	CTA	GCA	AAC	GAT	CCT	AGG	GAT	TTT	AAA	TCT	ACA	ATC	CCT	CAG	96
Lys	Ser	Leu	Ala	Asn	Asp	Pro	Arg	Asp	Phe	Lys	Ser	Thr	Ile	Pro	Gln	
			20					25					30			
AAC	GTC	AAC	CTG	TCT	GCA	GGA	TAC	TTA	GTT	ATT	AAA	GAG	GGG	GCC	GAA	144
Asn	Val	Asn	Leu	Ser	Ala	Gly	Tyr	Leu	Val	Ile	Lys	Glu	Gly	Ala	Glu	
		35					40					45				
GTC	ACA	GTT	TCA	AAA	TTC	ACG	CAG	TCT	CCA	GGA	TCG	CAT	TTA	GTT	TTA	192
Val	Thr	Val	Ser	Lys	Phe	Thr	Gln	Ser	Pro	Gly	Ser	His	Leu	Val	Leu	
	50					55					60					
GAT	TTA	GGA	ACC	AAA	CTG	ATA	ACC	TCT	AAG	GAA	GAC	ATT	GCC	ATC	ACA	240
Asp	Leu	Gly	Thr	Lys	Leu	Ile	Ala	Ser	Lys	Glu	Asp	Ile	Ala	Ile	Thr	
65					70				75					80		
GGC	CTC	GCG	ATA	GAT	ATA	GAT	AGC	TTA	AGC	TCA	TCC	TCA	ACA	GCA	GCT	288
Gly	Leu	Ala	Ile	Asp	Ile	Asp	Ser	Leu	Ser	Ser	Ser	Ser	Thr	Ala	Ala	
				85				90						95		

GTT ATT AAA GCA AAC ACC GCA AAT AAA CAG ATA TCC GTG ACG GAC TCT	336
Val Ile Lys Ala Asn Thr Ala Asn Lys Gln Ile Ser Val Thr Asp Ser	
100 105 110	
ATA GAA CTT ATC TCG CCT ACT GGC AAT GCC TAT GAA GAT CTC AGA ATG	384
Ile Glu Leu Ile Ser Pro Thr Gly Asn Ala Tyr Glu Asp Leu Arg Met	
115 120 125	
AGA AAT TCA CAG ACG TTC CCT CTG CTC TCT TTA GAG CCT GGA GCC GGG	432
Arg Asn Ser Gln Thr Phe Pro Leu Leu Ser Leu Glu Pro Gly Ala Gly	
130 135 140	
GGT AGT GTG ACT GTA ACT GCT GGA GAT TTC CTA CCG GTA AGT CCC CAT	480
Gly Ser Val Thr Val Thr Ala Gly Asp Phe Leu Pro Val Ser Pro His	
145 150 155 160	
TAT GGT TTT CAA GGC AAT TGG AAA TTA GCT TGG ACA GGA ACT GGA AAC	528
Tyr Gly Phe Gln Gly Asn Trp Lys Leu Ala Trp Thr Gly Thr Gly Asn	
165 170 175	
AAA GTT GGA GAA TTC TTC TGG GAT AAA ATA AAT TAT AAG CCT AGA CCT	576
Lys Val Gly Glu Phe Phe Trp Asp Lys Ile Asn Tyr Lys Pro Arg Pro	
180 185 190	
GAA AAA GAA GGA AAT TTA GTT CCT AAT ATC TTG TGG GGG AAT GCT GTA	624
Glu Lys Glu Gly Asn Leu Val Pro Asn Ile Leu Trp Gly Asn Ala Val	
195 200 205	
AAT GTC AGA TCC TTA ATG CAG GTT CAA GAG ACC CAT GCA TCG AGC TTA	672
Asn Val Arg Ser Leu Met Gln Val Gln Glu Thr His Ala Ser Ser Leu	
210 215 220	
CAG ACA GAT CGA GGG CTG TGG ATC GAT GGA ATT GGG AAT TTC TTC CAT	720
Gln Thr Asp Arg Gly Leu Trp Ile Asp Gly Ile Gly Asn Phe Phe His	
225 230 235 240	
GTA TCT GCC TCC GAA GAC AAT ATA AGG TAC CGT CAT AAC AGC GGT GGA	768
Val Ser Ala Ser Glu Asp Asn Ile Arg Tyr Arg His Asn Ser Gly Gly	
245 250 255	
TAT GTT CTA TCT GTA AAT AAT GAG ATC ACA CCT AAG CAC TAT ACT TCG	816
Tyr Val Leu Ser Val Asn Asn Glu Ile Thr Pro Lys His Tyr Thr Ser	
260 265 270	
ATG GCA TTT TCC CAA CTC TTT AGT AGA GAC AAA GAC TAT GCG GTT TCC	864
Met Ala Phe Ser Gln Leu Phe Ser Arg Asp Lys Asp Tyr Ala Val Ser	
275 280 285	
AAC AAC GAA TAC AGA ATG TAT TTA GGA TCG TAT CTC TAT CAA TAT ACA	912
Asn Asn Glu Tyr Arg Met Tyr Leu Gly Ser Tyr Leu Tyr Gln Tyr Thr	
290 295 300	
ACC TCC CTA GGG AAT ATT TTC CGT TAT GCT TCG CGT AAC CCT AAT GTA	960
Thr Ser Leu Gly Asn Ile Phe Arg Tyr Ala Ser Arg Asn Pro Asn Val	
305 310 315 320	
AAC GTC GGG ATT CTC TCA AGA AGG TTT CTT CAA AAT CCT CTT ATG ATT	1008

Asn Val Gly Ile Leu Ser Arg Arg Phe Leu Gln Asn Pro Leu Met Ile	
325 330 335	
TTT CAT TTT TTG TGT GCT TAT GGT CAT GCC ACC AAT GAT ATG AAA ACA	1056
Phe His Phe Leu Cys Ala Tyr Gly His Ala Thr Asn Asp Met Lys Thr	
340 345 350	
GAC TAC GCA AAT TTC CCT ATG GTG AAA AAC AGC TGG AGA AAC AAT TGT	1104
Asp Tyr Ala Asn Phe Pro Met Val Lys Asn Ser Trp Arg Asn Asn Cys	
355 360 365	
TGG GCT ATA AAA TGC GGA GGG AGC ATG CCT CTA TTG GTA TTT GAA AAC	1152
Trp Ala Ile Lys Cys Gly Gly Ser Met Pro Leu Leu Val Phe Glu Asn	
370 375 380	
GGA AAA CTT TTC CAA GGT GCC ATC CCA TTT ATG AAA CTA CAA TTA GTT	1200
Gly Lys Leu Phe Gln Gly Ala Ile Pro Phe Met Lys Leu Gln Leu Val	
385 390 395 400	

## (2) INFORMATION FOR SEQ ID NO:28:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 400 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: protein

## (v) FRAGMENT TYPE: internal

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:28:

Asp Pro Lys Asn Lys Glu Tyr Thr Gly Thr Ile Leu Phe Ser Gly Glu	
1 5 10 15	
Lys Ser Leu Ala Asn Asp Pro Arg Asp Phe Lys Ser Thr Ile Pro Gln	
20 25 30	
Asn Val Asn Leu Ser Ala Gly Tyr Leu Val Ile Lys Glu Gly Ala Glu	
35 40 45	
Val Thr Val Ser Lys Phe Thr Gln Ser Pro Gly Ser His Leu Val Leu	
50 55 60	
Asp Leu Gly Thr Lys Leu Ile Ala Ser Lys Glu Asp Ile Ala Ile Thr	
65 70 75 80	
Gly Leu Ala Ile Asp Ile Asp Ser Leu Ser Ser Ser Ser Thr Ala Ala	
85 90 95	
Val Ile Lys Ala Asn Thr Ala Asn Lys Gln Ile Ser Val Thr Asp Ser	
100 105 110	
Ile Glu Leu Ile Ser Pro Thr Gly Asn Ala Tyr Glu Asp Leu Arg Met	
115 120 125	
Arg Asn Ser Gln Thr Phe Pro Leu Leu Ser Leu Glu Pro Gly Ala Gly	
130 135 140	
Gly Ser Val Thr Val Thr Ala Gly Asp Phe Leu Pro Val Ser Pro His	
145 150 155 160	
Tyr Gly Phe Gln Gly Asn Trp Lys Leu Ala Trp Thr Gly Thr Gly Asn	
165 170 175	
Lys Val Gly Glu Phe Phe Trp Asp Lys Ile Asn Tyr Lys Pro Arg Pro	
180 185 190	

Glu Lys Glu Gly Asn Leu Val Pro Asn Ile Leu Trp Gly Asn Ala Val  
 195 200 205  
 Asn Val Arg Ser Leu Met Gln Val Gln Glu Thr His Ala Ser Ser Leu  
 210 215 220  
 Gln Thr Asp Arg Gly Leu Trp Ile Asp Gly Ile Gly Asn Phe Phe His  
 225 230 235 240  
 Val Ser Ala Ser Glu Asp Asn Ile Arg Tyr Arg His Asn Ser Gly Gly  
 245 250 255  
 Tyr Val Leu Ser Val Asn Asn Glu Ile Thr Pro Lys His Tyr Thr Ser  
 260 265 270  
 Met Ala Phe Ser Gln Leu Phe Ser Arg Asp Lys Asp Tyr Ala Val Ser  
 275 280 285  
 Asn Asn Glu Tyr Arg Met Tyr Leu Gly Ser Tyr Leu Tyr Gln Tyr Thr  
 290 295 300  
 Thr Ser Leu Gly Asn Ile Phe Arg Tyr Ala Ser Arg Asn Pro Asn Val  
 305 310 315 320  
 Asn Val Gly Ile Leu Ser Arg Arg Phe Leu Gln Asn Pro Leu Met Ile  
 325 330 335  
 Phe His Phe Leu Cys Ala Tyr Gly His Ala Thr Asn Asp Met Lys Thr  
 340 345 350  
 Asp Tyr Ala Asn Phe Pro Met Val Lys Asn Ser Trp Arg Asn Asn Cys  
 355 360 365  
 Trp Ala Ile Lys Cys Gly Gly Ser Met Pro Leu Leu Val Phe Glu Asn  
 370 375 380  
 Gly Lys Leu Phe Gln Gly Ala Ile Pro Phe Met Lys Leu Gln Leu Val  
 385 390 395 400

## (2) INFORMATION FOR SEQ ID NO:29:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1830 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: cDNA

## (ix) FEATURE:

- (A) NAME/KEY: Coding Sequence
- (B) LOCATION: 1...1830
- (D) OTHER INFORMATION:

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:29:

GAT CTC ACA TTA GGG AGT CGT GAC AGT TAT AAT GGT GAT ACA AGC ACC	48
Asp Leu Thr Leu Gly Ser Arg Asp Ser Tyr Asn Gly Asp Thr Ser Thr	
1 5 10 15	
ACA GAA TTT ACT CCT AAA GCG GCA ACT TCT GAT GCT AGT GGC ACG ACC	96
Thr Glu Phe Thr Pro Lys Ala Ala Thr Ser Asp Ala Ser Gly Thr Thr	
20 25 30	
TAT ATT CTC GAT GGG GAT GTC TCG ATA AGC CAA GCA GGG AAA CAA ACG	144
Tyr Ile Leu Asp Gly Asp Val Ser Ile Ser Gln Ala Gly Lys Gln Thr	
35 40 45	



AGC	TTA	ACC	ACA	AGT	TGT	TTT	TCT	AAC	ACT	GCA	GGA	AAT	CTT	ACC	TTC	192
Ser	Leu	Thr	Thr	Ser	Cys	Phe	Ser	Asn	Thr	Ala	Gly	Asn	Leu	Thr	Phe	
50						55					60					
TTA	GGG	AAC	GGA	TTT	TCT	CTT	CAT	TTT	GAC	AAT	ATT	ATT	TCG	TCT	ACT	240
Leu	Gly	Asn	Gly	Phe	Ser	Leu	His	Phe	Asp	Asn	Ile	Ile	Ser	Ser	Thr	
65					70				75						80	
GTT	GCA	GGT	GTT	GTT	GTT	AGC	AAT	ACA	GCA	GCT	TCT	GGG	ATT	ACG	AAA	288
Val	Ala	Gly	Val	Val	Val	Ser	Asn	Thr	Ala	Ala	Ser	Gly	Ile	Thr	Lys	
				85					90					95		
TTC	TCA	GGA	TTT	TCA	ACT	CTT	CGG	ATG	CTT	GCA	GCT	CCT	AGG	ACC	ACA	336
Phe	Ser	Gly	Phe	Ser	Thr	Leu	Arg	Met	Leu	Ala	Ala	Pro	Arg	Thr	Thr	
			100					105						110		
GGT	AAA	GGA	GCC	ATT	AAA	ATT	ACC	GAT	GGT	CTG	GTG	TTT	GAG	AGT	ATA	384
Gly	Lys	Gly	Ala	Ile	Lys	Ile	Thr	Asp	Gly	Leu	Val	Phe	Glu	Ser	Ile	
		115					120					125				
GGG	AAT	CTT	GAT	CCG	ATT	ACT	GTA	ACA	GGA	TCG	ACA	TCT	GTT	GCT	GAT	432
Gly	Asn	Leu	Asp	Pro	Ile	Thr	Val	Thr	Gly	Ser	Thr	Ser	Val	Ala	Asp	
	130					135					140					
GCT	CTC	AAT	ATT	AAT	AGC	CCT	GAT	ACT	GGA	GAT	AAC	AAA	GAG	TAT	ACG	480
Ala	Leu	Asn	Ile	Asn	Ser	Pro	Asp	Thr	Gly	Asp	Asn	Lys	Glu	Tyr	Thr	
145					150					155					160	
GGA	ACC	ATA	GTC	TTT	TCT	GGA	GAG	AAG	CTC	ACG	GAG	GCA	GAA	GCT	AAA	528
Gly	Thr	Ile	Val	Phe	Ser	Gly	Glu	Lys	Leu	Thr	Glu	Ala	Glu	Ala	Lys	
				165					170					175		
GAT	GAG	AAG	AAC	CGC	ACT	TCT	AAA	TTA	CTT	CAA	AAT	GTT	GCT	TTT	AAA	576
Asp	Glu	Lys	Asn	Arg	Thr	Ser	Lys	Leu	Leu	Gln	Asn	Val	Ala	Phe	Lys	
			180					185					190			
AAT	GGG	ACT	GTA	GTT	TTA	AAA	GGT	GAT	GTC	GTT	TTA	AGT	GCG	AAC	GGT	624
Asn	Gly	Thr	Val	Val	Leu	Lys	Gly	Asp	Val	Val	Leu	Ser	Ala	Asn	Gly	
		195					200						205			
TTC	TCT	CAG	GAT	GCA	AAC	TCT	AAG	TTG	ATT	ATG	GAT	TTA	GGG	ACG	TCG	672
Phe	Ser	Gln	Asp	Ala	Asn	Ser	Lys	Leu	Ile	Met	Asp	Leu	Gly	Thr	Ser	
	210					215					220					
TTG	GTT	GCA	AAC	ACC	GAA	AGT	ATC	GAG	TTA	ACG	AAT	TTG	GAA	ATT	AAT	720
Leu	Val	Ala	Asn	Thr	Glu	Ser	Ile	Glu	Leu	Thr	Asn	Leu	Glu	Ile	Asn	
225					230				235					240		
ATA	GAC	TCT	CTC	AGG	AAC	GGG	AAA	AAG	ATC	AAA	CTC	AGT	GCT	GCC	ACA	768
Ile	Asp	Ser	Leu	Arg	Asn	Gly	Lys	Lys	Ile	Lys	Leu	Ser	Ala	Ala	Thr	
				245					250					255		
GCT	CAG	AAA	GAT	ATT	CGT	ATA	GAT	CGT	CCT	GTT	GTA	CTG	GCA	ATT	AGC	816
Ala	Gln	Lys	Asp	Ile	Arg	Ile	Asp	Arg	Pro	Val	Val	Leu	Ala	Ile	Ser	
			260					265					270			
GAT	GAG	AGT	TTT	TAT	CAA	AAT	GGC	TTT	TTG	AAT	GAG	GAC	CAT	TCC	TAT	864

Asp	Glu	Ser	Phe	Tyr	Gln	Asn	Gly	Phe	Leu	Asn	Glu	Asp	His	Ser	Tyr	
	275						280					285				
GAT	GGG	ATT	CTT	GAG	TTA	GAT	GCT	GGG	AAA	GAC	ATC	GTG	ATT	TCT	GCA	912
Asp	Gly	Ile	Leu	Glu	Leu	Asp	Ala	Gly	Lys	Asp	Ile	Val	Ile	Ser	Ala	
	290					295					300					
GAT	TCT	CGC	AGT	ATA	GAT	GCT	GTA	CAA	TCT	CCG	TAT	GGC	TAT	CAG	GGA	960
Asp	Ser	Arg	Ser	Ile	Asp	Ala	Val	Gln	Ser	Pro	Tyr	Gly	Tyr	Gln	Gly	
305					310					315					320	
AAG	TGG	ACG	ATC	AAT	TGG	TCT	ACT	GAT	GAT	AAG	AAA	GCT	ACG	GTT	TCT	1008
Lys	Trp	Thr	Ile	Asn	Trp	Ser	Thr	Asp	Asp	Lys	Lys	Ala	Thr	Val	Ser	
			325					330						335		
TGG	GCG	AAG	CAG	AGT	TTT	AAT	CCC	ACT	GCT	GAG	CAG	GAG	GCT	CCG	TTA	1056
Trp	Ala	Lys	Gln	Ser	Phe	Asn	Pro	Thr	Ala	Glu	Gln	Glu	Ala	Pro	Leu	
			340					345					350			
GTT	CCT	AAT	CTT	CTT	TGG	GGT	TCT	TTT	ATA	GAT	GTT	CGT	TCC	TTC	CAG	1104
Val	Pro	Asn	Leu	Leu	Trp	Gly	Ser	Phe	Ile	Asp	Val	Arg	Ser	Phe	Gln	
		355					360					365				
AAT	TTT	ATA	GAG	CTA	GGT	ACT	GAA	GGT	GCT	CCT	TAC	GAA	AAG	AGA	TTT	1152
Asn	Phe	Ile	Glu	Leu	Gly	Thr	Glu	Gly	Ala	Pro	Tyr	Glu	Lys	Arg	Phe	
	370					375					380					
TGG	GTT	GCA	GGC	ATT	TCC	AAT	GTT	TTG	CAT	AGG	AGC	GGT	CGT	GAA	AAT	1200
Trp	Val	Ala	Gly	Ile	Ser	Asn	Val	Leu	His	Arg	Ser	Gly	Arg	Glu	Asn	
385					390					395					400	
CAA	AGG	AAA	TTC	CGT	CAT	GTG	AGT	GGA	GGT	GCT	GTA	GTA	GGT	GCT	AGC	1248
Gln	Arg	Lys	Phe	Arg	His	Val	Ser	Gly	Gly	Ala	Val	Val	Gly	Ala	Ser	
			405					410					415			
ACG	AGG	ATG	CCG	GGT	GGT	GAT	ACC	TTG	TCT	CTG	GGT	TTT	GCT	CAG	CTC	1296
Thr	Arg	Met	Pro	Gly	Gly	Asp	Thr	Leu	Ser	Leu	Gly	Phe	Ala	Gln	Leu	
			420					425					430			
TTT	GCG	CGT	GAC	AAA	GAC	TAC	TTT	ATG	AAT	ACC	AAT	TTC	GCA	AAG	ACC	1344
Phe	Ala	Arg	Asp	Lys	Asp	Tyr	Phe	Met	Asn	Thr	Asn	Phe	Ala	Lys	Thr	
		435					440					445				
TAC	GCA	GGA	TCT	TTA	CGT	TTG	CAG	CAC	GAT	GCT	TCC	CTA	TAC	TCT	GTG	1392
Tyr	Ala	Gly	Ser	Leu	Arg	Leu	Gln	His	Asp	Ala	Ser	Leu	Tyr	Ser	Val	
	450					455					460					
GTG	AGT	ATC	CTT	TTA	GGA	GAG	GGA	GGA	CTC	CGC	GAG	ATC	CTG	TTG	CCT	1440
Val	Ser	Ile	Leu	Leu	Gly	Glu	Gly	Gly	Leu	Arg	Glu	Ile	Leu	Leu	Pro	
465					470					475					480	
TAT	GTT	TCC	AAT	ACT	CTG	CCG	TGC	TCT	TTC	TAT	GGG	CAG	CTT	AGC	TAC	1488
Tyr	Val	Ser	Asn	Thr	Leu	Pro	Cys	Ser	Phe	Tyr	Gly	Gln	Leu	Ser	Tyr	
			485					490						495		
GGC	CAT	ACG	GAT	CAT	CGC	ATG	AAG	ACC	GAG	TCT	CTA	CCC	CCC	CCC	CCC	1536
Gly	His	Thr	Asp	His	Arg	Met	Lys	Thr	Glu	Ser	Leu	Pro	Pro	Pro	Pro	

500	505	510	
CCG ACG CTC TCG ACG GAT CAT ACT TCT TGG GGA GGA TAT GTC TGG GCT			1584
Pro Thr Leu Ser Thr Asp His Thr Ser Trp Gly Gly Tyr Val Trp Ala			
515	520	525	
GGA GAG CTG GGA ACT CGA GTT GCT GTT GAA AAT ACC AGC GGC AGA GGA			1632
Gly Glu Leu Gly Thr Arg Val Ala Val Glu Asn Thr Ser Gly Arg Gly			
530	535	540	
TTT TTC CGA GAG TAC ACT CCA TTT GTA AAA GTC CAA GCT GTT TAC TCG			1680
Phe Phe Arg Glu Tyr Thr Pro Phe Val Lys Val Gln Ala Val Tyr Ser			
545	550	555	560
CGC CAA GAT AGC TTT GTT GAA CTA GGA GCT ATC AGT CGT GAT TTT AGT			1728
Arg Gln Asp Ser Phe Val Glu Leu Gly Ala Ile Ser Arg Asp Phe Ser			
565	570	575	
GAT TCG CAT CTT TAT AAC CTT GCG ATT CCT CTT GGA ATC AAG TTA GAG			1776
Asp Ser His Leu Tyr Asn Leu Ala Ile Pro Leu Gly Ile Lys Leu Glu			
580	585	590	
AAA CGG TTT GCA GAG CAA TAT TAT CAT GTT GTT GCG ATG TAT TCT CCA			1824
Lys Arg Phe Ala Glu Gln Tyr Tyr His Val Val Ala Met Tyr Ser Pro			
595	600	605	
GAT GTT			1830
Asp Val			
610			

## (2) INFORMATION FOR SEQ ID NO:30:

## (i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 610 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

## (ii) MOLECULE TYPE: protein

## (v) FRAGMENT TYPE: internal

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:

```

Asp Leu Thr Leu Gly Ser Arg Asp Ser Tyr Asn Gly Asp Thr Ser Thr
 1           5           10           15
Thr Glu Phe Thr Pro Lys Ala Ala Thr Ser Asp Ala Ser Gly Thr Thr
      20           25           30
Tyr Ile Leu Asp Glu Asp Val Ser Ile Ser Gln Ala Gly Lys Gln Thr
      35           40           45
Ser Leu Thr Thr Ser Cys Phe Ser Asn Thr Ala Gly Asn Leu Thr Phe
      50           55           60
Leu Gly Asn Gly Phe Ser Leu His Phe Asp Asn Ile Ile Ser Ser Thr
      65           70           75           80
Val Ala Gly Val Val Val Ser Asn Thr Ala Ala Ser Gly Ile Thr Lys
      85           90           95
Phe Ser Gly Phe Ser Thr Leu Arg Met Leu Ala Ala Pro Arg Thr Thr

```

100 105 110  
 Gly Lys Gly Ala Ile Lys Ile Thr Asp Gly Leu Val Phe Glu Ser Ile  
 115 120 125  
 Gly Asn Leu Asp Pro Ile Thr Val Thr Gly Ser Thr Ser Val Ala Asp  
 130 135 140  
 Ala Leu Asn Ile Asn Ser Pro Asp Thr Gly Asp Asn Lys Glu Tyr Thr  
 145 150 155 160  
 Gly Thr Ile Val Phe Ser Gly Glu Lys Leu Thr Glu Ala Glu Ala Lys  
 165 170 175  
 Asp Glu Lys Asn Arg Thr Ser Lys Leu Leu Gln Asn Val Ala Phe Lys  
 180 185 190  
 Asn Gly Thr Val Val Leu Lys Gly Asp Val Val Leu Ser Ala Asn Gly  
 195 200 205  
 Phe Ser Gln Asp Ala Asn Ser Lys Leu Ile Met Asp Leu Gly Thr Ser  
 210 215 220  
 Leu Val Ala Asn Thr Glu Ser Ile Glu Leu Thr Asn Leu Glu Ile Asn  
 225 230 235 240  
 Ile Asp Ser Leu Arg Asn Gly Lys Lys Ile Lys Leu Ser Ala Ala Thr  
 245 250 255  
 Ala Gln Lys Asp Ile Arg Ile Asp Arg Pro Val Val Leu Ala Ile Ser  
 260 265 270  
 Asp Glu Ser Phe Tyr Gln Asn Gly Phe Leu Asn Glu Asp His Ser Tyr  
 275 280 285  
 Asp Gly Ile Leu Glu Leu Asp Ala Gly Lys Asp Ile Val Ile Ser Ala  
 290 295 300  
 Asp Ser Arg Ser Ile Asp Ala Val Gln Ser Pro Tyr Gly Tyr Gln Gly  
 305 310 315 320  
 Lys Trp Thr Ile Asn Trp Ser Thr Asp Asp Lys Lys Ala Thr Val Ser  
 325 330 335  
 Trp Ala Lys Gln Ser Phe Asn Pro Thr Ala Glu Gln Glu Ala Pro Leu  
 340 345 350  
 Val Pro Asn Leu Leu Trp Gly Ser Phe Ile Asp Val Arg Ser Phe Gln  
 355 360 365  
 Asn Phe Ile Glu Leu Gly Thr Glu Gly Ala Pro Tyr Glu Lys Arg Phe  
 370 375 380  
 Trp Val Ala Gly Ile Ser Asn Val Leu His Arg Ser Gly Arg Glu Asn  
 385 390 395 400  
 Gln Arg Lys Phe Arg His Val Ser Gly Gly Ala Val Val Gly Ala Ser  
 405 410 415  
 Thr Arg Met Pro Gly Gly Asp Thr Leu Ser Leu Gly Phe Ala Gln Leu  
 420 425 430  
 Phe Ala Arg Asp Lys Asp Tyr Phe Met Asn Thr Asn Phe Ala Lys Thr  
 435 440 445  
 Tyr Ala Gly Ser Leu Arg Leu Gln His Asp Ala Ser Leu Tyr Ser Val  
 450 455 460  
 Val Ser Ile Leu Leu Gly Glu Gly Gly Leu Arg Glu Ile Leu Leu Pro  
 465 470 475 480  
 Tyr Val Ser Asn Thr Leu Pro Cys Ser Phe Tyr Gly Gln Leu Ser Tyr  
 485 490 495  
 Gly His Thr Asp His Arg Met Lys Thr Glu Ser Leu Pro Pro Pro Pro  
 500 505 510  
 Pro Thr Leu Ser Thr Asp His Thr Ser Trp Gly Gly Tyr Val Trp Ala  
 515 520 525  
 Gly Glu Leu Gly Thr Arg Val Ala Val Glu Asn Thr Ser Gly Arg Gly  
 530 535 540  
 Phe Phe Arg Glu Tyr Thr Pro Phe Val Lys Val Gln Ala Val Tyr Ser  
 545 550 555 560

[illegible]